=> fil reg FILE 'REGISTRY' ENTERED AT 14:24:50 ON 30 SEP 2004 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2004 American Chemical Society (ACS)

Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 29 SEP 2004 HIGHEST RN 754169-63-6 DICTIONARY FILE UPDATES: 29 SEP 2004 HIGHEST RN 754169-63-6

TSCA INFORMATION NOW CURRENT THROUGH MAY 21, 2004

Please note that search-term pricing does apply when conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. For more information enter HELP PROP at an arrow prompt in the file or refer to the file summary sheet on the web at: http://www.cas.org/ONLINE/DBSS/registryss.html

=> d 185 ide can tot

L85 ANSWER 1 OF 4 REGISTRY COPYRIGHT 2004 ACS on STN

RN 81623-30-5 REGISTRY

CN 2(3H)-Furanone, dihydro-4-[(R)-hydroxy(4-hydroxy-3-methoxyphenyl)methyl]-3[(4-hydroxy-3-methoxyphenyl)methyl]-, (3R,4R)- (9CI) (CA INDEX NAME)
OTHER CA INDEX NAMES:

CN 2(3H)-Furanone, dihydro-4-[hydroxy(4-hydroxy-3-methoxyphenyl)methyl]-3-[(4-hydroxy-3-methoxyphenyl)methyl]-, [3R-[3 α ,4 β (R*)]]-OTHER NAMES:

CN (-)-allo-Hydroxymatairesinol

CN 5-Allohydroxymatairesinol

CN Allohydroxymatairesinol

FS STEREOSEARCH

MF C20 H22 O7

CI COM

LC STN Files: BEILSTEIN*, CA, CAPLUS, CASREACT, CHEMCATS, TOXCENTER, USPATFULL

(*File contains numerically searchable property data)

DT.CA CAplus document type: Journal; Patent

RL.P Roles from patents: BIOL (Biological study); RACT (Reactant or reagent); USES (Uses)

RL.NP Roles from non-patents: ANST (Analytical study); BIOL (Biological study); OCCU (Occurrence); PREP (Preparation); PROC (Process); PRP (Properties); RACT (Reactant or reagent); USES (Uses)

Absolute stereochemistry. Rotation (-).

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

21 REFERENCES IN FILE CA (1907 TO DATE)
21 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 141:156949

REFERENCE 2: 141:6955

REFERENCE 3: 139:197289

REFERENCE 4: 139:128028

REFERENCE 5: 139:117269

REFERENCE 6: 139:117267

REFERENCE 7: 139:8300

REFERENCE 8: 138:4458

REFERENCE 9: 135:166155

REFERENCE 10: 133:235125

L85 ANSWER 2 OF 4 REGISTRY COPYRIGHT 2004 ACS on STN

RN **78473-71-9** REGISTRY

CN 2(3H)-Furanone, dihydro-3,4-bis[(3-hydroxyphenyl)methyl]-, (3R,4R)-rel- (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN 2(3H)-Furanone, dihydro-3,4-bis[(3-hydroxyphenyl)methyl]-, trans-OTHER NAMES:

CN (±)-enterolactone

CN Enterolactone

CN HPMF

CN trans-2,3-Bis (3-hydroxybenzyl)- γ -butyrolactone

FS STEREOSEARCH

DR 76721-88-5, 82580-69-6, 110872-76-9

MF C18 H18 O4

CI COM

LC STN Files: AGRICOLA, ANABSTR, BEILSTEIN*, BIOBUSINESS, BIOSIS,
BIOTECHNO, CA, CAPLUS, CASREACT, CHEMCATS, CSCHEM, DDFU, DRUGU, EMBASE,
IPA, MRCK*, NAPRALERT, PROMT, TOXCENTER, USPAT2, USPATFULL
(*File contains numerically searchable property data)

DT.CA CAplus document type: Conference; Dissertation; Journal; Patent

RL.P Roles from patents: ANST (Analytical study); BIOL (Biological study); FORM (Formation, nonpreparative); OCCU (Occurrence); PREP (Preparation); PROC (Process); RACT (Reactant or reagent); USES (Uses)

RLD.P Roles for non-specific derivatives from patents: BIOL (Biological study); PREP (Preparation); USES (Uses)

RL.NP Roles from non-patents: ANST (Analytical study); BIOL (Biological study); FORM (Formation, nonpreparative); OCCU (Occurrence); PREP (Preparation); PROC (Process); PRP (Properties); RACT (Reactant or reagent); USES (Uses)

RLD.NP Roles for non-specific derivatives from non-patents: BIOL (Biological study); FORM (Formation, nonpreparative); OCCU (Occurrence); PROC (Process)

Relative stereochemistry.

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

225 REFERENCES IN FILE CA (1907 TO DATE)
6 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
228 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 141:222809

REFERENCE 2: 141:212883

REFERENCE 3: 141:206960

REFERENCE 4: 141:105811

REFERENCE 5: 141:104518

REFERENCE 6: 141:103196

REFERENCE 7: 141:99221

REFERENCE 8: 141:53197

REFERENCE 9: 140:420066

REFERENCE 10: 140:399547

L85 ANSWER 3 OF 4 REGISTRY COPYRIGHT 2004 ACS on STN

RN 20268-71-7 REGISTRY

CN 2(3H)-Furanone, dihydro-4-[(S)-hydroxy(4-hydroxy-3-methoxyphenyl)methyl]-3-[(4-hydroxy-3-methoxyphenyl)methyl]-, (3R,4R)- (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN 2(3H)-Furanone, dihydro-4-(α -hydroxyvanillyl)-3-vanillyl- (8CI)

CN 2 (3H) -Furanone, dihydro-4-[hydroxy(4-hydroxy-3-methoxyphenyl)methyl]-3-[(4-hydroxy-3-methoxyphenyl)methyl]-, [3R-[3α,4β(S*)]]-

OTHER NAMES:

CN (-)-Hydroxymatairesinol

CN α -Hydroxymatairesinol

CN 5-Hydroxymatairesinol

CN Hydroxymatairesinol

FS STEREOSEARCH

DR 29764-17-8

MF C20 H22 O7

CI COM

LC STN Files: AGRICOLA, ANABSTR, BEILSTEIN*, BIOSIS, CA, CAPLUS, CASREACT, CHEMCATS, IPA, NAPRALERT, PIRA, TOXCENTER, USPATZ, USPATFULL

(*File contains numerically searchable property data)

DT.CA CAplus document type: Conference; Journal; Patent

RL.P Roles from patents: BIOL (Biological study); OCCU (Occurrence); PREP (Preparation); PROC (Process); RACT (Reactant or reagent); USES (Uses)

RLD.P Roles for non-specific derivatives from patents: BIOL (Biological study); PREP (Preparation); USES (Uses)

RL.NP Roles from non-patents: ANST (Analytical study); BIOL (Biological study); OCCU (Occurrence); PREP (Preparation); PROC (Process); PRP (Properties); RACT (Reactant or reagent); USES (Uses)

Absolute stereochemistry. Rotation (-).

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

65 REFERENCES IN FILE CA (1907 TO DATE)

3 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

65 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 141:156949

REFERENCE 2: 141:99221

REFERENCE 3: 141:6955

REFERENCE 4: 140:219654

REFERENCE 5: 140:110360

REFERENCE 6: 140:65203

REFERENCE 7: 139:197289

REFERENCE 8: 139:179927

REFERENCE 9: 139:179926

REFERENCE 10: 139:128028

L85 ANSWER 4 OF 4 REGISTRY COPYRIGHT 2004 ACS on STN

RN 580-72-3 REGISTRY

CN 2(3H)-Furanone, dihydro-3,4-bis[(4-hydroxy-3-methoxyphenyl)methyl]-, (3R,4R)- (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN 2(3H)-Furanone, dihydro-3,4-bis[(4-hydroxy-3-methoxyphenyl)methyl]-,
(3R-trans)-

CN 2(3H)-Furanone, dihydro-3,4-divanillyl- (8CI)

CN Matairesinol (6CI)

OTHER NAMES:

CN (-)-Matairesinol

CN (8R,8'R)-(-)-Matairesinol

FS STEREOSEARCH

DR 41328-88-5

MF C20 H22 O6

CI COM

LC STN Files: AGRICOLA, ANABSTR, BEILSTEIN*, BIOBUSINESS, BIOSIS, BIOTECHNO, CA, CANCERLIT, CAOLD, CAPLUS, CASREACT, CHEMCATS, CSCHEM, DDFU, DRUGU, EMBASE, IPA, MEDLINE, NAPRALERT, PIRA, PROMT, SPECINFO, TOXCENTER, USPAT2, USPATFULL

(*File contains numerically searchable property data)

DT.CA CAplus document type: Conference; Journal; Patent

RL.P Roles from patents: BIOL (Biological study); OCCU (Occurrence); PREP (Preparation); PRP (Properties); RACT (Reactant or reagent); USES (Uses)

RLD.P Roles for non-specific derivatives from patents: BIOL (Biological study); PREP (Preparation); USES (Uses)

RL.NP Roles from non-patents: ANST (Analytical study); BIOL (Biological study); FORM (Formation, nonpreparative); MSC (Miscellaneous); OCCU (Occurrence); PREP (Preparation); PROC (Process); PRP (Properties); RACT (Reactant or reagent); USES (Uses); NORL (No role in record)

RLD.NP Roles for non-specific derivatives from non-patents: BIOL (Biological study); OCCU (Occurrence)

Absolute stereochemistry. Rotation (-).

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

283 REFERENCES IN FILE CA (1907 TO DATE)

6 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

286 REFERENCES IN FILE CAPLUS (1907 TO DATE)

6 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

REFERENCE 1: 141:206960

REFERENCE 2: 141:139670

REFERENCE 3: 141:139321

REFERENCE 4: 141:105811

REFERENCE 5: 141:99221

REFERENCE 6: 141:53096

REFERENCE 7: 141:6955

REFERENCE 8: 141:6381

REFERENCE 9: 140:411264

REFERENCE 10: 140:399547

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ANSWER 1 OF 2 REGISTRY COPYRIGHT 2004 ACS on STN
L86
    9003-99-0 REGISTRY
RN
CN
    Peroxidase (9CI) (CA INDEX NAME)
OTHER NAMES:
    Baylase RP
CN
    Biobake soy
CN
    Biobake Wheat
CN
CN
    Coniferyl alcohol peroxidase
    E.C. 1.11.1.7
CN
CN
    Enzylon OL 50
CN
    Eosinophil peroxidase
CN
    Extensin peroxidase
    Guaiacol peroxidase
CN
    Guaiacolase
CN
    Heme peroxidase
CN
    Lactoperoxidase
CN
    Manganese-dependent peroxidase
CN
CN
    Mn-dependent peroxidase
CN
    Myeloperoxidase
CN
    Novozym 502
CN
    Oxyperoxidase
CN
CN
    PEO-131
CN
    Peroxidase 51004
CN
     Protoheme peroxidase
CN
    Pyrocatechol peroxidase
CN
     Pyrogallol peroxidase
CN
    Scavengase p20
CN
     Scopoletin peroxidase
CN
     SP 502
    Thiocyanate peroxidase
CN
    Thiol peroxidase
CN
CN
     Verdoperoxidase
     9013-92-7, 9039-19-4, 191289-36-8
DR
    Unspecified
MF
    COM, MAN
CI
    STN Files:
                  ADISNEWS, AGRICOLA, ANABSTR, BIOBUSINESS, BIOSIS, BIOTECHNO,
LC
       CA, CAPLUS, CASREACT, CBNB, CEN, CHEMCATS, CHEMINFORMRX, CHEMLIST, CIN,
       CSCHEM, CSNB, DDFU, DRUGU, EMBASE, IFICDB, IFIPAT, IFIUDB, IPA, MEDLINE,
       MSDS-OHS, NAPRALERT, NIOSHTIC, PIRA, PROMT, TOXCENTER, ULIDAT, USPAT2,
       USPATFULL
                      EINECS**, TSCA**
    Other Sources:
         (**Enter CHEMLIST File for up-to-date regulatory information)
DT.CA CAplus document type: Book; Conference; Dissertation; Journal; Patent;
       Preprint; Report
RL.P
       Roles from patents: ANST (Analytical study); BIOL (Biological study);
       FORM (Formation, nonpreparative); MSC (Miscellaneous); OCCU
       (Occurrence); PREP (Preparation); PROC (Process); PRP (Properties); RACT
       (Reactant or reagent); USES (Uses); NORL (No role in record)
       Roles for non-specific derivatives from patents: ANST (Analytical
       study); BIOL (Biological study); CMBI (Combinatorial study); OCCU
       (Occurrence); PREP (Preparation); PROC (Process); PRP (Properties); RACT
       (Reactant or reagent); USES (Uses)
       Roles from non-patents: ANST (Analytical study); BIOL (Biological
       study); FORM (Formation, nonpreparative); MSC (Miscellaneous); OCCU
       (Occurrence); PREP (Preparation); PROC (Process); PRP (Properties); RACT
       (Reactant or reagent); USES (Uses); NORL (No role in record)
RLD.NP Roles for non-specific derivatives from non-patents: ANST (Analytical
       study); BIOL (Biological study); FORM (Formation, nonpreparative); MSC
       (Miscellaneous); OCCU (Occurrence); PREP (Preparation); PROC (Process);
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PRP (Properties); RACT (Reactant or reagent); USES (Uses)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

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36557 REFERENCES IN FILE CA (1907 TO DATE)
            2318 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
           36631 REFERENCES IN FILE CAPLUS (1907 TO DATE)
REFERENCE
            1: 141:230712
REFERENCE
            2:
                141:230109
REFERENCE
            3:
                141:225960
REFERENCE
                141:224708
REFERENCE
            5:
                141:224317
                141:224282
REFERENCE
            6:
REFERENCE
            7:
                141:224278
REFERENCE
            8: 141:224172
REFERENCE
            9:
                141:223221
REFERENCE 10: 141:223005
L86 ANSWER 2 OF 2 REGISTRY COPYRIGHT 2004 ACS on STN
     7782-44-7 REGISTRY
RN
     Oxygen (8CI, 9CI)
CN
                       (CA INDEX NAME)
OTHER NAMES:
CN
     Dioxygen
CN
     Molecular oxygen
CN
     Oxygen molecule
FS
     3D CONCORD
     1338-93-8, 14797-70-7, 80217-98-7, 80937-33-3
DR
MF
     02
CI
     COM
                  ADISNEWS, AGRICOLA, ANABSTR, BIOBUSINESS, BIOSIS, BIOTECHNO,
LC
       CA, CABA, CANCERLIT, CAPLUS, CASREACT, CBNB, CEN, CHEMCATS,
       CHEMINFORMRX, CHEMLIST, CHEMSAFE, CIN, CSCHEM, CSNB, DDFU, DETHERM*,
       DIOGENES, DIPPR*, DRUGU, EMBASE, ENCOMPLIT, ENCOMPLIT2, ENCOMPPAT,
       ENCOMPPAT2, GMELIN*, HSDB*, IFICDB, IFIPAT, IFIUDB, IPA, MEDLINE, MRCK*,
       MSDS-OHS, NIOSHTIC, PDLCOM*, PIRA, PROMT, PS, RTECS*, SPECINFO,
       TOXCENTER, TULSA, ULIDAT, USAN, USPATZ, USPATFULL, VTB
         (*File contains numerically searchable property data)
                     DSL**, EINECS**, TSCA**
         (**Enter CHEMLIST File for up-to-date regulatory information)
DT.CA
      CAplus document type: Book; Conference; Dissertation; Journal; Patent;
       Preprint; Report
RL.P
       Roles from patents: ANST (Analytical study); BIOL (Biological study);
       CMBI (Combinatorial study); FORM (Formation, nonpreparative); MSC
       (Miscellaneous); OCCU (Occurrence); PREP (Preparation); PROC (Process);
       PRP (Properties); RACT (Reactant or reagent); USES (Uses); NORL (No role
       in record)
RLD.P
      Roles for non-specific derivatives from patents: ANST (Analytical
       study); BIOL (Biological study); FORM (Formation, nonpreparative); MSC
       (Miscellaneous); OCCU (Occurrence); PREP (Preparation); PROC (Process);
       PRP (Properties); RACT (Reactant or reagent); USES (Uses)
      Roles from non-patents: ANST (Analytical study); BIOL (Biological
RL.NP
       study); CMBI (Combinatorial study); FORM (Formation, nonpreparative);
       MSC (Miscellaneous); OCCU (Occurrence); PREP (Preparation); PROC
       (Process); PRP (Properties); RACT (Reactant or reagent); USES (Uses);
       NORL (No role in record)
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RLD.NP Roles for non-specific derivatives from non-patents: ANST (Analytical

study); BIOL (Biological study); CMBI (Combinatorial study); FORM
(Formation, nonpreparative); MSC (Miscellaneous); OCCU (Occurrence);
PREP (Preparation); PROC (Process); PRP (Properties); RACT (Reactant or reagent); USES (Uses)

o = 0

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

349327 REFERENCES IN FILE CA (1907 TO DATE)
27975 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
349909 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 141:235211 REFERENCE 2: 141:235146 REFERENCE 141:235138 REFERENCE 141:234918 REFERENCE 5: 141:234912 REFERENCE 141:234833 6: REFERENCE 7: 141:234796 REFERENCE 8: 141:234727 REFERENCE 9: 141:234726

REFERENCE 10: 141:234500

=> d his

(FILE 'HOME' ENTERED AT 13:25:41 ON 30 SEP 2004) SET COST OFF

FILE 'HCAPLUS' ENTERED AT 13:26:09 ON 30 SEP 2004 L1 1 S US20030100514/PN OR US2001-991971#/AP,PRN

FILE 'REGISTRY' ENTERED AT 13:26:42 ON 30 SEP 2004 L23 S 580-72-3 OR 20268-71-7 OR 78473-71-9 E C20H2207/MF E C20H22O7/MF L337 S E3 AND 46.150.18/RID AND OC4/ES AND 3/NR L4 26 S L3 AND 3 METHOXY L5 26 S L4 AND 4 HYDROXY L6 21 S L5 AND FURANONE SEL RN 1 6 7 8 10 11 16 20 L7 8 S E1-E8 rs7 S L7 NOT 718614-97-2 SEL RN 4 5 1.9 5 S L8 NOT E9-E10 L10 32 S L3 NOT L9 E C20H22O6/MF 56 S E3 AND OC4/ES AND 46.150.18/RID AND 3/NR L11 12 S L11 AND 4 HYDROXY AND 3 METHOXY AND FURANONE L124 S L12 NOT (D/ELS OR 13C# OR LABELED) L13

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E C18H18O4/MF
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L14
              8 S L14 AND 3 HYDROXY AND BIS AND FURANONE
L15
              5 S L15 NOT (D/ELS OR LABELED OR 13C)
L16
             14 S L2, L9, L13, L16
L17
              9 S L17 AND (?MATAIRESINOL? OR ?ENTEROLACTON?)/CNS
L18
              5 S L17 NOT L18
L19
             14 S L17-L19
L20
                SEL RN
              8 S E1-E14/CRN
L21
     FILE 'HCAPLUS' ENTERED AT 13:39:29 ON 30 SEP 2004
            527 S L20
L22
            583 S ENTEROLACTON? OR HYDROXYMATAIRESINOL? OR MATAIRESINOL?
L23
            640 S L22, L23
L24
                E AHOTUPA M/AU
             91 S E3-E5
L25
                E ERIKSSON J/AU
L26
            221 S E3-E11, E34-E36
                E KANGAS L/AU
            127 S E3-E5, E8-E11
L27
                E UNKILA M/AU
L28
             48 S E3-E5
                E KOMI J/AU
             12 S E3-E6
L29
                E PERALA M/AU
             21 S E3, E4, E6
L30
                E KORTE H/AU
             23 S E3, E4, E10
L31
                E HORMOS/PA,CS
             27 S E3-E19
L32
             16 S L24 AND L25-L32
L33
                E PHAGOCYTE/CT
           3427 S E3,E12
L34
                E E12+ALL
                E E2+ALL
          32274 S E5+NT
L35
                E NEUTROPHIL/CT
                E E3+ALL
          29239 S E24,E23
L36
                E T CELL/CT
                E E4+ALL
L37
          40180 S E20-E23
          70058 S E19+NT
L38
                E E18+ALL
L39
         169033 S E19, E18+NT
                E MYELOID/CT
                E E11+ALL
           2697 S E2
L40
              4 S L24 AND L34-L40
L41
                E ANIMAL RESPIRATION/CT
           1613 S E3 (L) BURST
L42
                E RESPIRATION, ANIMAL/CT
L43
           1421 S E4
                E REACTIVE OXYGEN/CT
                E E4+ALL
L44
          22365 S E3
              2 S L24 AND L42-L44
L45
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FILE 'HCAPLUS' ENTERED AT 14:09:52 ON 30 SEP 2004

1 S OXYGEN/CN

L46

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L47
              1 S L24 AND L46
                E LIGNAN/CT
                E E4+ALL
            356 S L24 AND E2
L48
L49
            356 S L24 AND E2+NT
L50
              4 S L41, L45, L47
              1 S L50 AND L33
L51
L52
              3 S L50 AND L48, L49
              4 S L50-L52
L53
             65 S L20 (L) (THU OR DMA OR PAC OR PKT)/RL
L54
            160 S L24 AND (PHARMACEUT? OR PHARMACOL? OR PATHOL? OR IMMUN?)/SC,S
L55
            164 S L54, L55
L56
              9 S L56 AND L33
L57
              3 S L56 AND L53
L58
              4 S L53, L58
L59
              8 S L57 NOT L59
L60
     FILE 'REGISTRY' ENTERED AT 14:15:46 ON 30 SEP 2004
L61
              1 S 9003-99-0
     FILE 'HCAPLUS' ENTERED AT 14:15:57 ON 30 SEP 2004
              2 S L61 AND L24
L62
L63
              1 S L24 AND MYELOPEROXIDASE
              6 S L24 AND ?PEROXIDASE?
L64
              6 S L62-L64
L65
             17 S L59, L60, L65
L66
             14 S L66 AND (PD<=20011126 OR PRD<=20011126 OR AD<=20011126)
L67
              3 S L66 NOT L67
L68
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              8 S L67 NOT E1-E18
L69
             11 S L68, L69
L70
                E TRANPLANTATION/CT
                E TRANSPLANTATION/CT
            812 S E3
L71
                E TRANSPLANT/CT
            494 S E3
L72
          87407 S E5+OLD, NT, PFT, RT
L73
L74
           5085 S E61
          76222 S E69+OLD, NT, PFT, RT
L75
            812 S E72, E74
L76
                E E3+ALL
                E E2+ALL
L77
           7719 S E7-E16
L78
          35079 S E6+NT
           6674 S E43+NT
L79
L80
          30585 S E42+NT
              5 S L24 AND L71-L80
L81
L82
              4 S L81 NOT AROMATASE/TI
             14 S L70, L82 AND L1, L22-L45, L47-L60, L62-L82
L83
                SEL HIT RN
     FILE 'REGISTRY' ENTERED AT 14:24:16 ON 30 SEP 2004
L84
              6 S E1-E6
L85
              4 S L84 AND L20
L86
              2 S L84 AND L61,L46
     FILE 'REGISTRY' ENTERED AT 14:24:50 ON 30 SEP 2004
=> fil hcaplus
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=> fil hcaplus

FILE 'HCAPLUS' ENTERED AT 14:25:17 ON 30 SEP 2004

USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.

PLEASE SEE "HELP USAGETERMS" FOR DETAILS.

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FILE COVERS 1907 - 30 Sep 2004 VOL 141 ISS 14 FILE LAST UPDATED: 29 Sep 2004 (20040929/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

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=> d 183 all hitstr tot
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L83 ANSWER 1 OF 14 HCAPLUS COPYRIGHT 2004 ACS on STN

AN 2004:738451 HCAPLUS

ED Entered STN: 10 Sep 2004

TI Lignans and neolignans useful as cathepsin inhibitors, and their use

IN Jean, Daniel; Rabhi, Cherif; Schwaab, Veronique

PA LMD, Fr.

SO Fr. Demande, 24 pp.

CODEN: FRXXBL
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DT Patent

LA French

IC ICM A61K031-357 ICS A61K031-343; C07D319-20; C07D307-80; C07D407-10; A61P035-00; C07D319-00; C07D307-00

CC 1-12 (Pharmacology)

FAN.CNT 1

	PA	PATENT NO.						DATE		i	APPLICATION NO.						DATE			
	PI FR	FR 2851919			A1 20040910			0910	Ī	FR 2	003-	20030303								
	WO	WO 2004080379					A2 20040923				WO 2	004-	20040303							
		W:	ΑE,	AG,	AL,	AM,	ΑT,	AU,	AZ,	BA,	BB,	BG,	BR,	BW,	BY,	ΒZ,	CA,	CH,		
			CN,	CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	EG,	ES,	FI,	GB,	GD,		
			GE,	GH,	GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	KP,	KR,	KZ,	LC,		
			LK,	LR,	LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NA,	NI,		
			NO,	NZ,	OM,	PG,	PH,	PL,	PT,	RO,	RU,	SC,	SD,	SE,	SG,	SK,	SL,	SY,		
			ТJ,	TM,	TN,	TR,	TT,	TZ,	UA,	UG,	US,	UZ,	VC,	VN,	YU,	ZA,	ZM,	ZW		
		RW	: BW,	GH,	GM,	KE,	LS,	MW,	MZ,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ZW,	AM,	AZ,		
			BY,	KG,	KZ,	MD,	RU,	ТJ,	TM,	AT,	BE,	BG,	CH,	CY,	CZ,	DE,	DK,	EE,		
																	SE,	-		
		•	SK,	TR,	BF,	ВJ,	CF,	CG,	CI,	CM,	GA,	GN,	GQ,	GW,	ML,	MR,	NE,	SN,		
			TD,	TG			·	-								•	·	·		
PRAI FR 2003-2584				Α	A 20030303															
	CLASS																			

DATEN

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PATENT NO. CLASS PATENT FAMILY CLASSIFICATION CODES

FR 2851919 ICM A61K031-357
ICS A61K031-343; C07D319-20; C07D307-80; C07D407-10; A61P035-00; C07D319-00; C07D307-00
```

- AB The invention discloses the use of lignans and neolignans as inhibitors of cathepsins. The compds. of the invention may be used e.g. to inhibit metastases and as hepatoprotectants.
- ST cathepsin inhibitor lignan neolignan; metastasis inhibition hepatoprotection lignan cathepsin inhibitor

```
IT
     INDEXING IN PROGRESS
IT
     Cytoprotective agents
         (hepatoprotective; lignan and neolignan cathepsin inhibitors, and use)
IT
     Toxicity
         (hepatotoxicity; lignan and neolignan cathepsin inhibitors, and use)
IT
     Antitumor agents
     Cosmetics
     Feed
     Fish
     Food
     Human
       Organ preservation
     Pangium edule
         (lignan and neolignan cathepsin inhibitors, and use)
IT
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
         (lignan and neolignan cathepsin inhibitors, and use)
IT
     Lignans
     Natural products, pharmaceutical
       Neolignans
     RL: COS (Cosmetic use); FFD (Food or feed use); PAC (Pharmacological
     activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
         (lignan and neolignan cathepsin inhibitors, and use)
IT
     Neoplasm
        (metastasis; lignan and neolignan cathepsin inhibitors, and use)
IT
     Liver
        (toxicity; lignan and neolignan cathepsin inhibitors, and use)
     9004-08-4, Cathepsin 9047-22-7, Cathepsin B
TΤ
                                                      60616-82-2, Cathepsin L
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (lignan and neolignan cathepsin inhibitors, and use)
IT
     77053-44-2P, Americanin B
                                 77053-45-3P, Americanin D
                                                              133838-66-1P,
     Isoamericanol A
                       214344-43-1P
     RL: COS (Cosmetic use); FFD (Food or feed use); NPO (Natural product
     occurrence); PAC (Pharmacological activity); PUR (Purification or
     recovery); THU (Therapeutic use); BIOL (Biological study); OCCU
     (Occurrence); PREP (Preparation); USES (Uses)
        (lignan and neolignan cathepsin inhibitors, and use)
TΤ
     580-72-3, Matairesinol
                              29388-59-8,
                            59332-00-2, Eusiderin
     Secoisolariciresinol
     RL: COS (Cosmetic use); FFD (Food or feed use); PAC (Pharmacological
     activity); THU (Therapeutic use); BIOL (Biological study);
     USES (Uses)
        (lignan and neolignan cathepsin inhibitors, and use)
              THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT
(1) Cassidy, A; WO 02080702 A 2002 HCAPLUS
(2) Eckerman, C; WO 0059946 A 2000 HCAPLUS
(3) Ford, J; PLANT POLYPHENOLS 2: CHEMISTRY, BIOLOGY, PHARMACOLOGY, ECOLOGY
    1999, P675 HCAPLUS
(4) Gu, W; TETRAHEDRON LETTERS 2000, V41(32), P6079 HCAPLUS
(5) Madaus & Co Dr; GB 2035300 A 1980 HCAPLUS
(6) Matsumoto, K; TETRAHEDRON LETTERS 1999, V40(16), P3185 HCAPLUS
(7) Merck & Co Inc; EP 0159565 A 1985 HCAPLUS
(8) Sampath, K; US 6489514 B1 2002 HCAPLUS
TT
     INDEXING IN PROGRESS
IT
     580-72-3, Matairesinol
     RL: COS (Cosmetic use); FFD (Food or feed use); PAC (Pharmacological
     activity); THU (Therapeutic use); BIOL (Biological study);
     USES (Uses)
        (lignan and neolignan cathepsin inhibitors, and use)
RN
     580-72-3 HCAPLUS
CN
     2(3H)-Furanone, dihydro-3,4-bis[(4-hydroxy-3-methoxyphenyl)methyl]-,
```

(3R,4R) - (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

OMe

OH

OMe

0=

HO

IT

Drugs

use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(cyclodextrin complexes; lignan complexes with cyclodextrin and their uses in food products, dietary supplements or pharmaceutical compns.)

Health products

(lignan complexes with cyclodextrin and their uses in food products, dietary supplements or pharmaceutical compns.)

Inclusion compounds IT

IT

RL: FFD (Food or feed use); IMF (Industrial manufacture); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(lignan complexes with cyclodextrin and their uses in food products,

dietary supplements or pharmaceutical compns.)

484-39-9DP, Liovil, lignan derivs., cyclodextrin complexes Pinoresinol, lignan derivs., cyclodextrin complexes α -Conidendrin, lignan derivs., cyclodextrin complexes 548-29-8DP, Isolariciresinol, lignan derivs., cyclodextrin complexes 580-72-3DP, Matairesinol, lignan derivs., cyclodextrin 1177-14-6DP, DL-Syringaresinol, lignan derivs., cyclodextrin 7585-39-9DP, β -Cyclodextrin, hydroxypropyl ether, complexes inclusion compds. with lignans 7585-39-9DP, β -Cyclodextrin, 7770-78-7DP, Arctigenin, lignan derivs., inclusion compds. with lignans 10016-20-3DP, α -Cyclodextrin, inclusion cyclodextrin complexes compds. with lignans 17465-86-0DP, γ -Cyclodextrin, inclusion compds. with lignans 20268-71-7DP, Hydroxymatairesinol lignan derivs., cyclodextrin complexes 27003-73-2DP, Lariciresinol, lignan derivs., cyclodextrin complexes 29388-59-8DP, Secoisolariciresinol, lignan derivs., cyclodextrin complexes 34444-37-6DP, Nortrachelogenin, lignan derivs., cyclodextrin complexes

53250-61-6DP, Oxomatairesinol, lignan derivs., cyclodextrin complexes 568593-01-1DP, Picearesinol, lignan derivs., cyclodextrin complexes RL: FFD (Food or feed use); IMF (Industrial manufacture); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(lignan complexes with cyclodextrin and their uses in food products, dietary supplements or pharmaceutical compns.)

THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD RE.CNT RE

(1) Suntory Limited; EP 0387000 A2 1990 HCAPLUS

- (2) van Uden, W; J Nat Prod 1997, V60, P401 HCAPLUS
- (3) Vincieri, F; Il Farmaco 1994, V49(1), P63 HCAPLUS
- 580-72-3DP, Matairesinol, lignan derivs., cyclodextrin complexes 20268-71-7DP, Hydroxymatairesinol, lignan derivs., cyclodextrin complexes RL: FFD (Food or feed use); IMF (Industrial manufacture); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES

(lignan complexes with cyclodextrin and their uses in food products, dietary supplements or pharmaceutical compns.)

RN580-72-3 HCAPLUS

2(3H)-Furanone, dihydro-3,4-bis[(4-hydroxy-3-methoxyphenyl)methyl]-, CN(CA INDEX NAME) (3R, 4R) - (9CI)

Absolute stereochemistry. Rotation (-).

20268-71-7 HCAPLUS RN

2(3H)-Furanone, dihydro-4-[(S)-hydroxy(4-hydroxy-3-methoxyphenyl)methyl]-3-CN [(4-hydroxy-3-methoxyphenyl)methyl]-, (3R,4R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

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ANSWER 3 OF 14 HCAPLUS COPYRIGHT 2004 ACS on STN
L83
    2004:2687 HCAPLUS
AN
    140:65203
DN
    Entered STN: 02 Jan 2004
ED
TI
    Lignan topical formulations
    Korte, Helena; Lehtola, Veli-Matti; Unkila, Mikko;
IN
    Hiilovaara-Teijo, Mervi; Ahotupa, Markku
    Hormos Nutraceutical Oy Ltd., Finland
PA
    PCT Int. Appl., 25 pp.
SO
    CODEN: PIXXD2
DT
    Patent
LA
    English
    ICM A61K031-341
IC
CC
    63-6 (Pharmaceuticals)
    Section cross-reference(s): 62
FAN.CNT 1
                                      APPLICATION NO.
                             DATE
    PATENT NO.
                       KIND
                              _____
                                         ______
                       _ _ _ _
     ______
                                        WO 2003-FI375
                       A1
                              20031231
                                                               20030515
    WO 2004000304
PI
        W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
            CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
            GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
            LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
            PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ,
            UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD,
            RU, TJ, TM
        RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG,
            CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC,
            NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ,
            GW, ML, MR, NE, SN, TD, TG
    FI 2002001184
                        Α
                              20031220
                                          FI 2002-1184
                                                                20020619
                              20020619
PRAI FI 2002-1184
                        Α
CLASS
 PATENT NO.
                CLASS PATENT FAMILY CLASSIFICATION CODES
                      A61K031-341
WO 2004000304 ICM
    MARPAT 140:65203
OS
GI
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CN

Absolute stereochemistry. Rotation (-).

This invention concerns a topical formulation comprising a lignan or AB lignan ester in a dermatol. acceptable vehicle. The formulation can be either a cosmetic formulation or a pharmaceutical formulation. E.g., water-in-oil emulsions contained a lignan such as hydroxymatairesinol (I) or matairesinol dibutyrate, an emulsifier such as sorbitan fatty acid ester, humectant such as glycerol, preservative, and water. STlignan topical pharmaceutical cosmetic ITAntioxidants Cosmetics (lignan topical formulations) ITLignans RL: COS (Cosmetic use); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (lignan topical formulations) ITDrug delivery systems (topical; lignan topical formulations) 484-39-9, Liovil 487-36-5, Pinoresinol 518-55-8, α -Conidendrin IT 7770-78-7, Arctigenin 548-29-8, Isolariciresinol 1177-14-6 27003-73-2, 20268-71-7, Hydroxymatairesinol Lariciresinol 29388-59-8, Secoisolariciresinol 34444-37-6, Nortrachelogenin 53250-61-6, Oxomatairesinol 568593-01-1, Picearesinol RL: COS (Cosmetic use); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (lignan topical formulations) RE.CNT THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD RE(1) Cedars-Sinai Medical Center; WO 0103687 A2 2001 HCAPLUS (2) Unilever N V; WO 0108651 A1 2001 HCAPLUS 20268-71-7, Hydroxymatairesinol ITRL: COS (Cosmetic use); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (lignan topical formulations) 20268-71-7 HCAPLUS RN

2(3H)-Furanone, dihydro-4-[(S)-hydroxy(4-hydroxy-3-methoxyphenyl)methyl]-3-

[(4-hydroxy-3-methoxyphenyl)methyl]-, (3R,4R)- (9CI) (CA INDEX NAME)

IT

Prostate gland, disease

patients)

```
ANSWER 4 OF 14 HCAPLUS COPYRIGHT 2004 ACS on STN
L83
     2003:590821 HCAPLUS
AN
     139:128028
DN
     Entered STN: 01 Aug 2003
ED
     Method for prevention of diseases in coeliac patients
TI
IN
     Unkila, Mikko
     Finland
PΑ
     U.S. Pat. Appl. Publ., 5 pp.
SO
     CODEN: USXXCO
DT
     Patent
LA
     English
     ICM A61K031-70
IC
     ICS A61K035-78
     514022000; 514025000; 424769000
NCL
CC
     1-9 (Pharmacology)
FAN.CNT 1
                                          APPLICATION NO.
                        KIND
                                                                 DATE
                               DATE
     PATENT NO.
     ______
                                           _____
                        ----
                               _____
                               20030731
                                           US 2002-54900
                                                                 20020125
                         A1
PI
     US 2003144216
                        A1
                               20030731
                                           WO 2003-FI6
                                                                 20030107
     WO 2003061649
         W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
            CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
            GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
            LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
            PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ,
            UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD,
            RU, TJ, TM
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG,
            CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC,
            NL, PT, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW,
            ML, MR, NE, SN, TD, TG
PRAI US 2002-54900
                         Α
                               20020125
CLASS
                CLASS PATENT FAMILY CLASSIFICATION CODES
 PATENT NO.
                       ______
 US 2003144216
                ICM
                       A61K031-70
                       A61K035-78
                ICS
                       514022000; 514025000; 424769000
                NCL
     Methods for prevention of cancers, precancers, certain non-cancer, hormone
AB
     dependent diseases and/or cardiovascular diseases in a person suffering
     from coeliac disease, based on administering of a lignan to the person. A
     method for increasing the level of enterolactone or another
     metabolite of a lignan in a person's serum is also disclosed, where the
     person suffers from coeliac disease, thereby causing prevention of a
     cancer or a certain non-cancer, hormone dependent disease in the person,
     based on administering of a lignan to the person.
     celiac disease prevention treatment
ST
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(benign hyperplasia; method for prevention of diseases in celiac

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IT
     Intestine, neoplasm
        (colon; method for prevention of diseases in celiac patients)
IT
     Urethra
        (dyssynergia; method for prevention of diseases in celiac patients)
IT
     Intestine, neoplasm
        (familial polyposis; method for prevention of diseases in celiac
        patients)
IT
     Mammary gland, disease
        (gynecomastia, in men; method for prevention of diseases in celiac
        patients)
IT
     Lipoproteins
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (low-d., oxidized; method for prevention of diseases in celiac
        patients)
IT
     Urinary tract
        (lower, disease; method for prevention of diseases in celiac patients)
IT
     Cardiovascular agents
     Celiac disease
     Esophagus, neoplasm
     Human
     Mammary gland, neoplasm
     Neoplasm
     Prostate gland, neoplasm
     Testis, neoplasm
        (method for prevention of diseases in celiac patients)
IT
     Lignans
     RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (plant; method for prevention of diseases in celiac patients)
     Intestine, neoplasm
ΙT
        (small; method for prevention of diseases in celiac patients)
                                               518-55-8, Conidendrin
                       487-36-5, Pinoresinol
TΤ
     484-39-9, Liovil
     548-29-8, Isolariciresinol 580-72-3, Matairesinol
     7770-78-7, (-)-Arctigenin 11041-15-9, Conidendric acid
     20268-71-7, Hydroxymatairesinol
                                       27003-73-2,
                   29388-59-8, Secoisolariciresinol
     Lariciresinol
                                                         34444-37-6.
                       41607-20-9 53250-61-6, Oxomatairesinol
     Nortrachelogenin
     81623-30-5, Allohydroxymatairesinol
                                           84413-77-4, (+)-Arctigenin
     568593-01-1, Picearesinol
     RL: PAC (Pharmacological activity); THU (Therapeutic
     use); BIOL (Biological study); USES (Uses)
        (method for prevention of diseases in celiac patients)
IT
     580-72-3, Matairesinol 20268-71-7,
     Hydroxymatairesinol 81623-30-5, Allohydroxymatairesinol
     RL: PAC (Pharmacological activity); THU (Therapeutic
     use); BIOL (Biological study); USES (Uses)
        (method for prevention of diseases in celiac patients)
RN
     580-72-3 HCAPLUS
     2(3H)-Furanone, dihydro-3,4-bis[(4-hydroxy-3-methoxyphenyl)methyl]-,
CN
                    (CA INDEX NAME)
     (3R, 4R) - (9CI)
```

Absolute stereochemistry. Rotation (-).

RN 20268-71-7 HCAPLUS

CN 2(3H)-Furanone, dihydro-4-[(S)-hydroxy(4-hydroxy-3-methoxyphenyl)methyl]-3-[(4-hydroxy-3-methoxyphenyl)methyl]-, (3R,4R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

RN 81623-30-5 HCAPLUS

CN 2 (3H) -Furanone, dihydro-4-[(R)-hydroxy(4-hydroxy-3-methoxyphenyl)methyl]-3-[(4-hydroxy-3-methoxyphenyl)methyl]-, (3R,4R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

L83 ANSWER 5 OF 14 HCAPLUS COPYRIGHT 2004 ACS on STN

AN 2003:414080 HCAPLUS

DN 138:379228

ED Entered STN: 30 May 2003

TI Method using lignans for inhibiting overactivity of phagocytes or lymphocytes in an individual, and therapeutic use

IN Ahotupa, Markku; Eriksson, John; Kangas, Lauri; Unkila, Mikko; Komi, Janne; Perala, Merja; Korte, Helena

PA Finland

SO U.S. Pat. Appl. Publ., 10 pp.

CODEN: USXXCO

DT Patent

LA English

IC ICM A61K031-365 ICS A61K031-05

NCL 514022000; 514460000; 514731000

CC 1-7 (Pharmacology)

FAN CNT 1

	FAN.	CNT 1											•				_				
	PATENT NO.							D	DATE			APPLICATION NO.						DATE			
						_															
	ΡI	PI US 2003100514					A 1		20030529			US 2001-991971						20011126 <			
		WO 2003045376							20030605			WO 2002-FI936					20021121 <				
		V	7 :	ΑE,	AG,	AL,	AM,	AT,	ΑU,	AZ,	BA,	BB,	BG,	BR,	BY,	BZ,	CA,	CH,	CN,		
				CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	ES,	FI,	GB,	GD,	GE,	GH,		
									IN,												
									MD,												
									SE,												
				UA.	UG.	US,	UZ,	VC,	VN,	YU,	ZA,	ZM,	ZW,	AM,	ΑZ,	BY,	KG,	KZ,	MD,		

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RU, TJ, TM
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG,
             CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR,
             NE, SN, TD, TG
PRAI US 2001-991971
                                 20011126 <--
CLASS
                 CLASS PATENT FAMILY CLASSIFICATION CODES
 PATENT NO.
                 ICM
                        A61K031-365
 US 2003100514
                 ICS
                        A61K031-05
                        514022000; 514460000; 514731000
                 NCL
     The invention provides a method for inhibiting the overactivity of
AB
     phagocytes or lymphocytes in an individual by administering to the
     individual an effective amount of a lignan, wherein (i) the phagocytes are
     neutrophils and the lignan is hydroxymatairesinol or
     matairesinol or mixts. thereof; or (ii) the phagocytes are cells
     of myeloid origin and the lignan is enterolactone or
     hydroxymatairesinol or mixts. thereof; or (iii) the lymphocytes
     are T-lymphocytes and the lignan is hydroxymatairesinol,
     matairesinol or enterolactone or mixts. thereof.
     invention also provides a method for treating or preventing an acute
     ischemia-reperfusion injury or a chronic condition, caused by overactivity
     of phagocytes or lymphocytes in an individual, the method comprising
     decreasing the activity of phagocytes in an individual by administering to
     the individual an effective amount of a lignan.
     lignan phagocyte lymphocye overactivity inhibition therapeutic; neutrophil
     overactivity inhibition hydroxymatairesinol matairesinol
     therapeutic; myeloid cell overactivity inhibition enterolactone
     hydroxymatairesinol therapeutic; T lymphocyte overactivity
     inhibition hydroxymatairesinol matairesinol
     enterolactone therapeutic; ischemia reperfusion injury therapeutic
     lignan
IT
     Intestine, disease
         (Crohn's; lignans for inhibiting overactivity of phagocytes or
        lymphocytes, and therapeutic use)
ΙT
     Apoptosis
        (Fas-induced; lignans for inhibiting overactivity of phagocytes or
        lymphocytes, and therapeutic use)
     Tumor necrosis factors
IT
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (TNF-\alpha, release; lignans for inhibiting overactivity of
        phagocytes or lymphocytes, and therapeutic use)
IT
     Respiratory distress syndrome
        (adult; lignans for inhibiting overactivity of phagocytes or
        lymphocytes, and therapeutic use)
IT
     Nervous system, disease
        (amyotrophic lateral sclerosis; lignans for inhibiting overactivity of
        phagocytes or lymphocytes, and therapeutic use)
IT
     Antiarteriosclerotics
        (antiatherosclerotics; lignans for inhibiting overactivity of
        phagocytes or lymphocytes, and therapeutic use)
IT
     Respiration, animal
        (burst; lignans for inhibiting overactivity of phagocytes or
        lymphocytes, and therapeutic use)
IT
     Drugs
        (gastrointestinal; lignans for inhibiting overactivity of phagocytes or
        lymphocytes, and therapeutic use)
     Shock (circulatory collapse)
IT
        (hemorrhagic; lignans for inhibiting overactivity of phagocytes or
        lymphocytes, and therapeutic use)
     Hypercholesterolemia
IT
         (hypercholesterolemic atherosclerosis; lignans for inhibiting
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overactivity of phagocytes or lymphocytes, and therapeutic use)
     Atherosclerosis
IT
        (hypercholesterolemic; lignans for inhibiting overactivity of
        phagocytes or lymphocytes, and therapeutic use)
TI
        (hypersensitivity, type I and type II; lignans for inhibiting
        overactivity of phagocytes or lymphocytes, and therapeutic use)
     Heart, disease
IT
        (infarction; lignans for inhibiting overactivity of phagocytes or
        lymphocytes, and therapeutic use)
     Intestine, disease
IT
        (inflammatory; lignans for inhibiting overactivity of phagocytes or
        lymphocytes, and therapeutic use)
     Reperfusion
IT
        (injury; lignans for inhibiting overactivity of phagocytes or
        lymphocytes, and therapeutic use)
     Diabetes mellitus
IT
        (insulin-dependent; lignans for inhibiting overactivity of phagocytes
        or lymphocytes, and therapeutic use)
     Heart, disease
ΤT
        (ischemia; lignans for inhibiting overactivity of phagocytes or
        lymphocytes, and therapeutic use)
IT
     AIDS (disease)
     Allergy
     Allergy inhibitors
     Alzheimer's disease
     Anti-AIDS agents
     Anti-Alzheimer's agents
     Anti-inflammatory agents
     Anti-ischemic agents
     Antiasthmatics
     Antidiabetic agents
     Antiparkinsonian agents
     Antirheumatic agents
     Antiviral agents
     Asthma
     Autoimmune disease
     Cardiovascular agents
     Cataract
     Dermatitis
     Human
     Human immunodeficiency virus
       Immunosuppressants
     Inflammation
     Ischemia
       Lymphocyte
     Monocyte
     Multiple sclerosis
       Neutrophil
     Osteoporosis
     Parkinson's disease
       Phagocyte
     Psoriasis
     Rheumatoid arthritis
       T cell (lymphocyte)
       Transplant and Transplantation
       Transplant rejection
        (lignans for inhibiting overactivity of phagocytes or lymphocytes, and
        therapeutic use)
     Fas antigen
IT
       Reactive oxygen species
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
```

(lignans for inhibiting overactivity of phagocytes or lymphocytes, and

```
therapeutic use)
IT
    Liquans
     RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (lignans for inhibiting overactivity of phagocytes or lymphocytes, and
        therapeutic use)
    Hematopoietic precursor cell
TT
        (myeloid, phagocyte of myeloid origin; lignans for
        inhibiting overactivity of phagocytes or lymphocytes, and therapeutic
        use)
    Diabetes mellitus
TΤ
        (non-insulin-dependent; lignans for inhibiting overactivity of
        phagocytes or lymphocytes, and therapeutic use)
     Shock (circulatory collapse)
TT
        (septic; lignans for inhibiting overactivity of phagocytes or
        lymphocytes, and therapeutic use)
     Brain, disease
TT
        (stroke; lignans for inhibiting overactivity of phagocytes or
        lymphocytes, and therapeutic use)
     7782-44-7D, Oxygen, reactive species 9003-99-0,
IT
     Myeloperoxidase
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (lignans for inhibiting overactivity of phagocytes or lymphocytes, and
        therapeutic use)
     580-72-3, Matairesinol 20268-71-7,
IT
     Hydroxymatairesinol 78473-71-9, Enterolactone
     RL: PAC (Pharmacological activity); THU (Therapeutic
     use); BIOL (Biological study); USES (Uses)
        (lignans for inhibiting overactivity of phagocytes or lymphocytes, and
        therapeutic use)
     7782-44-7D, Oxygen, reactive species 9003-99-0,
TT
     Myeloperoxidase
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (lignans for inhibiting overactivity of phagocytes or lymphocytes, and
        therapeutic use)
     7782-44-7 HCAPLUS
RN
     Oxygen (8CI, 9CI) (CA INDEX NAME)
CN
     9003-99-0 HCAPLUS
RN
     Peroxidase (9CI) (CA INDEX NAME)
CN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
     580-72-3, Matairesinol 20268-71-7,
IT
     Hydroxymatairesinol 78473-71-9, Enterolactone
     RL: PAC (Pharmacological activity); THU (Therapeutic
     use); BIOL (Biological study); USES (Uses)
        (lignans for inhibiting overactivity of phagocytes or lymphocytes, and
        therapeutic use)
     580-72-3 HCAPLUS
RN
     2(3H)-Furanone, dihydro-3,4-bis[(4-hydroxy-3-methoxyphenyl)methyl]-,
CN
     (3R,4R) - (9CI) (CA INDEX NAME)
```

Absolute stereochemistry. Rotation (-).

RN 20268-71-7 HCAPLUS

CN 2 (3H) -Furanone, dihydro-4-[(S)-hydroxy(4-hydroxy-3-methoxyphenyl)methyl]-3-[(4-hydroxy-3-methoxyphenyl)methyl]-, (3R,4R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

RN 78473-71-9 HCAPLUS

CN 2(3H)-Furanone, dihydro-3,4-bis[(3-hydroxyphenyl)methyl]-, (3R,4R)-rel-(9CI) (CA INDEX NAME)

Relative stereochemistry.

- L83 ANSWER 6 OF 14 HCAPLUS COPYRIGHT 2004 ACS on STN
- AN 2002:392225 HCAPLUS
- DN 136:380145
- ED Entered STN: 24 May 2002
- Prevention of cancers, non-cancerous hormone-dependent diseases, and cardiovascular diseases by use of hydroxymatairesinol, and a pharmaceutical preparation, food additive and food product comprising hydroxymatairesinol
- IN Ahotupa, Markku; Eckerman, Christer; Kangas, Lauri; Makela, Sari; Saarinen, Niina; Santti, Risto; Warri, Anni
- PA Hormos Nutraceutical oy Ltd., Finland
- SO U.S. Pat. Appl. Publ., 15 pp., Cont.-in-part of U.S. Ser. No. 829,944. CODEN: USXXCO
- DT Patent
- LA English
- IC ICM A61K031-70 ICS A61K035-78
- NCL 514022000
- CC 1-12 (Pharmacology)

Section cross-reference(s): 18, 63

FAN.CNT 2

PATENT NO.

KIND DATE

APPLICATION NO.

DATE

```
US 2002061854 EA1
                               20020523
                                           US 2001-972850
                                                                  20011010 <--
                         B2
     US 6689809
                               20040210
                        B1
                                           US 1999-281094
     US 6451849
                               20020917
                                                                  19990330 <--
                        A1
A1
     US 2001016590
                               20010823
                                          US 2001-829944
                                                                 20010411 <--
                                        US 2003-639530
     US 2004048804
                               20040311
                                                                 20030813 <--
PRAI US 1999-281094
                               19990330 <--
     US 2001-829944
                         A2
                               20010411 <--
     US 2001-972850
                         A1
                               20011010 <--
CLASS
                CLASS PATENT FAMILY CLASSIFICATION CODES
 PATENT NO.
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 US 2002061854
                 ICM
                       A61K031-70
                 ICS
                       A61K035-78
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                ECLA
 US 2002061854
                       A23L001/30B; A61K031/365
                                                                           <--
                ECLA
 US 6451849
                       A23L001/30B; A61K031/365
                                                                           <--
 US 2004048804
                ECLA
                       A23L001/30B; A61K031/365
AB
     The invention discloses methods for prevention of cancers, certain
     non-cancerous, hormone-dependent diseases, and/or cardiovascular diseases
     in a person, based on the administration of hydroxymatairesinol.
     The invention also discloses a method for increasing the level of
     enterolactone or another metabolite of hydroxymatairesinol
     in a person's serum, thereby causing prevention of a cancer or a certain
     non-cancerous, hormone-dependent disease in a person, based on
     administration of hydroxymatairesinol. Furthermore, the
     invention discloses pharmaceutical prepns., food additives, and food
     products comprising hydroxymatairesinol.
ST
     hydroxymatairesinol pharmaceutical food antitumor cardiovascular
     drug; hormone dependent disease pharmaceutical hydroxymatairesinol
      enterolactone stimulation therapeutic metabolite
     hydroxymatairesinol
IT
     Animal cell line
        (JEG-3; hydroxymatairesinol for prevention of cancers,
        non-cancerous hormone-dependent diseases, and cardiovascular diseases,
        and pharmaceutical and food products)
IT
    Animal cell line
        (MCF-7; hydroxymatairesinol for prevention of cancers,
       non-cancerous hormone-dependent diseases, and cardiovascular diseases,
       and pharmaceutical and food products)
\mathbf{IT}
    Health food
        (and designer foods; hydroxymatairesinol for prevention of
       cancers, non-cancerous hormone-dependent diseases, and cardiovascular
       diseases, and pharmaceutical and food products)
IT
    Drug delivery systems
        (and nutraceuticals; hydroxymatairesinol for prevention of
       cancers, non-cancerous hormone-dependent diseases, and cardiovascular
       diseases, and pharmaceutical and food products)
IT
        (and pharmafoods; hydroxymatairesinol for prevention of
       cancers, non-cancerous hormone-dependent diseases, and cardiovascular
       diseases, and pharmaceutical and food products)
TT
        (bran; hydroxymatairesinol for prevention of cancers,
       non-cancerous hormone-dependent diseases, and cardiovascular diseases,
       and pharmaceutical and food products)
IT
    Flaxseed
        (flour; hydroxymatairesinol for prevention of cancers,
       non-cancerous hormone-dependent diseases, and cardiovascular diseases,
       and pharmaceutical and food products)
```

IT Antioxidants
Daucus carota
Glycine max

IT

IT

TT

IT

IT

IT

TT

TT

IT

IT

IT

IT

Nutrients Onion (Allium cepa) Picea abies Secale cereale Solanum tuberosum Wheat bran (hydroxymatairesinol for prevention of cancers, non-cancerous hormone-dependent diseases, and cardiovascular diseases, and pharmaceutical and food products) Lignans RL: BSU (Biological study, unclassified); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (hydroxymatairesinol for prevention of cancers, non-cancerous hormone-dependent diseases, and cardiovascular diseases, and pharmaceutical and food products) Peroxidation (lipid; hydroxymatairesinol for prevention of cancers, non-cancerous hormone-dependent diseases, and cardiovascular diseases, and pharmaceutical and food products) Lipoproteins RL: BSU (Biological study, unclassified); BIOL (Biological study). (low-d., oxidation; hydroxymatairesinol for prevention of cancers, non-cancerous hormone-dependent diseases, and cardiovascular diseases, and pharmaceutical and food products) Antitumor agents (mammary gland; hydroxymatairesinol for prevention of cancers, non-cancerous hormone-dependent diseases, and cardiovascular diseases, and pharmaceutical and food products) Mammary gland (neoplasm, inhibitors; hydroxymatairesinol for prevention of cancers, non-cancerous hormone-dependent diseases, and cardiovascular diseases, and pharmaceutical and food products) Bran (oat; hydroxymatairesinol for prevention of cancers, non-cancerous hormone-dependent diseases, and cardiovascular diseases, and pharmaceutical and food products) Lipids, biological studies RL: BSU (Biological study, unclassified); BIOL (Biological study) (peroxidn.; hydroxymatairesinol for prevention of cancers, non-cancerous hormone-dependent diseases, and cardiovascular diseases, and pharmaceutical and food products) Peroxides, biological studies RL: BSU (Biological study, unclassified); BIOL (Biological study) (radicals; hydroxymatairesinol for prevention of cancers, non-cancerous hormone-dependent diseases, and cardiovascular diseases, and pharmaceutical and food products) Wood (soft; hydroxymatairesinol for prevention of cancers, non-cancerous hormone-dependent diseases, and cardiovascular diseases, and pharmaceutical and food products) Diet (supplements; hydroxymatairesinol for prevention of cancers, non-cancerous hormone-dependent diseases, and cardiovascular diseases, and pharmaceutical and food products) 9039-48-9, Aromatase 518-55-8, α -Conidendrin 11041-15-9, Conidendric acid 11062-77-4, Superoxide RL: BSU (Biological study, unclassified); BIOL (Biological study) (hydroxymatairesinol for prevention of cancers, non-cancerous hormone-dependent diseases, and cardiovascular diseases, and pharmaceutical and food products) 80226-00-2, Enterodiol 78473-71-9, Enterolactone

RL: BSU (Biological study, unclassified); PAC (Pharmacological

activity); BIOL (Biological study)

(hydroxymatairesinol for prevention of cancers, non-cancerous hormone-dependent diseases, and cardiovascular diseases, and pharmaceutical and food products)

IT 20268-71-7, Hydroxymatairesinol

RL: NPO (Natural product occurrence); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); OCCU (Occurrence); USES (Uses)

(hydroxymatairesinol for prevention of cancers, non-cancerous hormone-dependent diseases, and cardiovascular diseases, and pharmaceutical and food products)

IT 117-39-5, Quercetin 128-37-0, BHT, biological studies 491-54-3, Kaempferide 520-18-3, Kaempferol 25013-16-5, BHA 53188-07-1, Trolox 380448-80-6

RL: PAC (Pharmacological activity); BIOL (Biological study)
(hydroxymatairesinol for prevention of cancers, non-cancerous hormone-dependent diseases, and cardiovascular diseases, and pharmaceutical and food products)

IT 20268-71-7D, Hydroxymatairesinol, (stereo)isomers RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(hydroxymatairesinol for prevention of cancers, non-cancerous hormone-dependent diseases, and cardiovascular diseases, and pharmaceutical and food products)

IT 78473-71-9, Enterolactone

RL: BSU (Biological study, unclassified); PAC (Pharmacological activity); BIOL (Biological study)

(hydroxymatairesinol for prevention of cancers, non-cancerous hormone-dependent diseases, and cardiovascular diseases, and pharmaceutical and food products)

RN 78473-71-9 HCAPLUS

CN 2(3H)-Furanone, dihydro-3,4-bis[(3-hydroxyphenyl)methyl]-, (3R,4R)-rel-(9CI) (CA INDEX NAME)

Relative stereochemistry.

IT 20268-71-7, Hydroxymatairesinol

RL: NPO (Natural product occurrence); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); OCCU (Occurrence); USES (Uses)

(hydroxymatairesinol for prevention of cancers, non-cancerous hormone-dependent diseases, and cardiovascular diseases, and pharmaceutical and food products)

RN 20268-71-7 HCAPLUS

CN 2 (3H) -Furanone, dihydro-4-[(S)-hydroxy(4-hydroxy-3-methoxyphenyl)methyl]-3-[(4-hydroxy-3-methoxyphenyl)methyl]-, (3R,4R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

IT 20268-71-7D, Hydroxymatairesinol, (stereo)isomers RL: PAC (Pharmacological activity); THU (Therapeutic

use); BIOL (Biological study); USES (Uses)

(hydroxymatairesinol for prevention of cancers, non-cancerous hormone-dependent diseases, and cardiovascular diseases, and pharmaceutical and food products)

RN 20268-71-7 HCAPLUS

CN 2(3H)-Furanone, dihydro-4-[(S)-hydroxy(4-hydroxy-3-methoxyphenyl)methyl]-3-[(4-hydroxy-3-methoxyphenyl)methyl]-, (3R,4R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

L83 ANSWER 7 OF 14 HCAPLUS COPYRIGHT 2004 ACS on STN

AN 2001:922174 HCAPLUS

DN 136:291701

ED Entered STN: 21 Dec 2001

TI Immunosuppressive constituents from Saussurea medusa

AU Duan, Hongquan; Takaishi, Yoshihisa; Momota, Hiroshi; Ohmoto, Yasukazu; Taki, Takao

CS Faculty of Pharmaceutical Sciences, University of Tokushima, Tokushima, 770-8505, Japan

SO Phytochemistry (2002), 59(1), 85-90 CODEN: PYTCAS; ISSN: 0031-9422

PB Elsevier Science Ltd.

DT Journal

LA English

CC 11-1 (Plant Biochemistry) Section cross-reference(s): 26

GΙ

The methanol extract of Saussurea medusa Maxim afforded two lignans: (e.g. I) and 1-hydroxy-2,4-guaicyl-3,7-dioxabicyclo[3.3.0]octane; two chlorophyll derivs.: 13-epi-phaeophorbide-a and 13-epi-phaeophorbide-a Me ester; one megastigmane derivative: 3-hydroxy-5,6-epoxy-7-megastigmen-9-one (II), along with 19 known compds. Their structures were established on the basis of spectroscopic studies.

ST lignan chlorophyll megastigmane deriv Saussurea immunosuppressant

T

Chlorophylls, biological studies

RL: BSU (Biological study, unclassified); NPO (Natural product occurrence); PAC (Pharmacological activity); PRP (Properties); PUR (Purification or recovery); BIOL (Biological study); OCCU (Occurrence); PREP (Preparation)

(derivs.; immunosuppressive constituents from Saussurea medusa)

IT Immunosuppressants

Saussurea medusa

(immunosuppressive constituents from Saussurea medusa)

IT Lignans

IT

RL: BSU (Biological study, unclassified); NPO (Natural product occurrence); PAC (Pharmacological activity); PRP (Properties); PUR (Purification or recovery); BIOL (Biological study); OCCU (Occurrence); PREP (Preparation)

(immunosuppressive constituents from Saussurea medusa)

IT Cytokines

RL: BSU (Biological study, unclassified); BIOL (Biological study) (inhibition effect on cytokinins of immunosuppressive constituents from Saussurea medusa)

IT New natural products

(lignans, chlorophyll derivs. and megastigmane derivative from Saussurea medusa)

IT Molecular structure, natural product

(of lignans, chlorophyll derivs. and megastigmane derivative from Saussurea medusa)

IT 64070-09-3P, 13-epi-Phaeophorbide-a methyl ester 78964-31-5P, 13-epi-Phaeophorbide-a 175418-93-6P 408513-60-0P 408513-62-2P, 1α-Hydroxy-2α, 4α-guaicyl-3,7-dioxabicyclo[3.3.0]octane RL: BSU (Biological study, unclassified); NPO (Natural product occurrence); PAC (Pharmacological activity); PRP (Properties); PUR (Purification or recovery); BIOL (Biological study); OCCU (Occurrence); PREP (Preparation)

(immunosuppressive constituents from Saussurea medusa)

487-36-5, (+)-Pinoresinol **580-72-3**, **Matairesinol** TT 603-17-8, Pheophytin a 3147-18-0, Pheophytin b 5594-30-9, Methyl 5989-02-6, Loliolide 6216-81-5, Lirioresinol B phaeophoride a 20240-17-9 7770-78-7, Arctigenin 15664-29-6, Phaeophorbide a 27003-73-2, 24404-50-0, Epipinoresinol 20362-31-6, Arctiin 29388-59-8, Secoisolariciresinol 40957-99-1, Lariciresinol 126882-59-5, 79733-01-0 79733-03-2 99305-01-8 (+)-Medioresinol (-)-Berchemol RL: BSU (Biological study, unclassified); PAC (Pharmacological activity); BIOL (Biological study) (immunosuppressive constituents from Saussurea medusa) IT 408512-16-3P RL: PRP (Properties); SPN (Synthetic preparation); PREP (Preparation) (preparation and properties of) THERE ARE 16 CITED REFERENCES AVAILABLE FOR THIS RECORD RE.CNT RE (1) Briggs, L; Journal of Chemical Society (C) 1968, P3042 HCAPLUS (2) Chan, Y; Chemical and Pharmaceutical Bulletin 1999, V47, P887 HCAPLUS (3) Duan, H; Phytochemistry 2000, V53, P805 HCAPLUS (4) Fang, J; Phytochemistry 1989, V28, P3553 HCAPLUS (5) Fonseca, S; Phytochemistry 1978, V17, P499 (6) Hodges, R; Tetrahedron 1964, V20, P1463 HCAPLUS (7) Kita, M; Microbiology and Immunology 1992, V36, P507 HCAPLUS (8) Kobayashi, M; Chemical and Pharmaceutical Bulletin 1991, V39, P3348 HCAPLUS (9) Li, Y; Phytochemistry 1989, V28, P3395 HCAPLUS (10) Nakatani, Y; Chemical and Pharmaceutical Bulletin 1981, V29, P2261 HCAPLUS (11) Rahman, M; Phytochemistry 1990, V29, P1971 HCAPLUS (12) Sakurai, N; Chemical and Pharmaceutical Bulletin 1989, V37, P3311 HCAPLUS

HCAPLUS (15) Wray, V; Tetrahedron 1979, V35, P2275 HCAPLUS

(13) Takeda, Y; Phytochemistry 1997, V44, P1335 HCAPLUS

(16) Yang, R; Natural Medicines 1997, V51, P134

IT 580-72-3, Matairesinol

RL: BSU (Biological study, unclassified); PAC (Pharmacological activity); BIOL (Biological study)

(14) Tsukamoto, H; Chemical and Pharmaceutical Bulletin 1984, V32, P4482

(immunosuppressive constituents from Saussurea medusa)

RN 580-72-3 HCAPLUS

CN 2(3H)-Furanone, dihydro-3,4-bis[(4-hydroxy-3-methoxyphenyl)methyl]-, (3R,4R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

L83 ANSWER 8 OF 14 HCAPLUS COPYRIGHT 2004 ACS on STN

AN 2001:573545 HCAPLUS

DN 135:132430

ED Entered STN: 08 Aug 2001

TI Decreasing the intracellular level of β-catenin by administering hydroxymatairesinol, and therapeutic and diagnostic methods

'IN Mutanen, Marja

PA Hormos Nutraceutical Oy Ltd., Finland

SO U.S., 7 pp. CODEN: USXXAM

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DT
     Patent
LA
     English
IC
     ICM A61K031-00
NCL
     514461000
     1-6 (Pharmacology)
CC
     Section cross-reference(s): 9
FAN.CNT 1
     PATENT NO.
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                                               APPLICATION NO.
                                                                          DATE
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                                                                        20000417 <--
                                                US 2000-550602
PΙ
     US 6271257
                            B1
                                   20010807
                           A1
     WO 2001078720
                                   20011025
                                                WO 2001-FI110
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              SD, SE, SG
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
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                            A1
                                  20030409 EP 2001-905844
                                                                         20010208 <--
              AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR
                            T2
                                  20031014
                                              JP 2001-576020
                                                                          20010208 <--
PRAI US 2000-550602
                            Α
                                   20000417
     WO 2001-FI110
                           W
                                   20010208 <--
CLASS
                 CLASS PATENT FAMILY CLASSIFICATION CODES
 PATENT NO.
 _____
                          ______
                  ICM
 US 6271257
                          A61K031-00
                  NCL
                          514461000
     A method is provided for decreasing the intracellular, especially nuclear,
level
     of \beta\text{-catenin} in an individual. Also provided is a method for the
     prevention or treatment of a disease or condition in an individual,
     wherein the disease or condition is related to a mutant APC gene or to an
     elevated level of intracellular \beta-catenin. Specifically provided is
     a method for the treatment of familial adenomatous polyposis.
     Furthermore, the invention provides methods for screening a subject to
     determine if said subject is a carrier of a mutant APC gene, as well as methods
     for diagnosing an individual's predisposition for a disease or condition
     in an individual, the disease or condition being related to a mutant APC
     gene or to an elevated level of intracellular \beta-catenin.
     hydroxymatairesinol therapeutic beta catenin redn; APC gene
ST
     disease diagnosis therapy hydroxymatairesinol; familial
     adenomatous polyposis treatment hydroxymatairesinol
IT
     Gene, animal
     RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL
     (Biological study); PROC (Process)
         (APC; Decreasing the intracellular level of \beta-catenin by
        administering hydroxymatairesinol, and therapeutic and
        diagnostic methods)
     Antitumor agents
TT
     Mutation
     Rye
         (Decreasing the intracellular level of \beta-catenin by administering
        hydroxymatairesinol, and therapeutic and diagnostic methods)
IT
     Intestine, neoplasm
         (adenoma; Decreasing the intracellular level of \beta-catenin by
        administering hydroxymatairesinol, and therapeutic and
        diagnostic methods)
IT
     Intestine, neoplasm
         (familial polyposis; Decreasing the intracellular level of
```

 β -catenin by administering **hydroxymatairesinol**, and therapeutic and diagnostic methods)

IT Catenins

RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)

 $(\beta$ -; Decreasing the intracellular level of β -catenin by administering hydroxymatairesinol, and therapeutic and diagnostic methods)

IT 20268-71-7, Hydroxymatairesinol

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(Decreasing the intracellular level of β -catenin by administering hydroxymatairesinol, and therapeutic and diagnostic methods)

RE.CNT 7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD RE

(1) Anon; WO 9213103 1992 HCAPLUS

- (2) Barker; US 5998600 1999 HCAPLUS
- (3) Bras; European Journal of Cancer Prevention 1999, V8(4), P305 MEDLINE
- (4) Herter; Journal of Cancer Research and Clinical Oncology 1999, V125(5) HCAPLUS
- (5) Kinzler; US 5709998 1998 HCAPLUS
- (6) Mahmoud; Proceeding of the American Association for Cancer Research Annual Meeting 1999, V40, P530
- (7) Saarinen; Nutrition and Cancer 2000, V36(2) HCAPLUS
- IT 20268-71-7, Hydroxymatairesinol

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(Decreasing the intracellular level of β -catenin by administering hydroxymatairesinol, and therapeutic and diagnostic methods)

RN 20268-71-7 HCAPLUS

CN 2(3H)-Furanone, dihydro-4-[(S)-hydroxy(4-hydroxy-3-methoxyphenyl)methyl]-3-[(4-hydroxy-3-methoxyphenyl)methyl]-, (3R,4R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-):

- L83 ANSWER 9 OF 14 HCAPLUS COPYRIGHT 2004 ACS on STN
- AN 2001:450166 HCAPLUS
- DN 135:189741
- ED Entered STN: 22 Jun 2001
- TI Anti-AIDS Agents. 46. Anti-HIV Activity of Harman, an Anti-HIV Principle from Symplocos setchuensis, and Its Derivatives
- AU Ishida, Junko; Wang, Hui-Kang; Oyama, Masayoshi; Cosentino, Mark L.; Hu, Chang-Qi; Lee, Kuo-Hsiung
- CS Natural Products Laboratory School of Pharmacy, University of North Carolina, Chapel Hill, NC, 27599-7360, USA
- SO Journal of Natural Products (2001), 64(7), 958-960 CODEN: JNPRDF; ISSN: 0163-3864
- PB American Chemical Society
- DT Journal

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English
LA
     1-3 (Pharmacology)
CC
     Section cross-reference(s): 63
     Matairesinol and harman, identified from Symplocos setchuensis,
AB
     were found to inhibit HIV replication in H9 lymphocyte cells. Anti-HIV
     evaluation of 28 derivs. of harman revealed that compound 19 showed potent
     activity with EC50 and therapeutic index values of 0.037 µM and 210,
     antiHIV Symplocos harman deriv SAR
ST
IT
     Lymphocyte
        (H9; anti-HIV principle from Symplocos setchuensis, and its derivs.)
IT
     Anti-AIDS agents
     Structure-activity relationship
     Symplocos
        (anti-HIV principle from Symplocos setchuensis, and its derivs.)
     244-63-3, 9H-Pyrido[3,4-b]indole 442-51-3, Harmine 486-84-0, Harman
IT
                                      6415-92-5
                                                  6519-18-2
               525-41-7
                          6028-07-5
                                                               10593-56-3,
     9H-Pyrido[3,4-b]indole, 7-ethoxy-1-methyl-
                                                               24415-61-0
                                                  17019-08-8
                                             199530-62-6
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     85645-27-8
                 143502-37-8
                               186790-81-8
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     257938-78-6
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                                 356790-37-9
     257938-86-6
                   356790-36-8
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
        (anti-HIV principle from Symplocos setchuensis, and its derivs.)
              THERE ARE 11 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT
        11
RE
(1) Abe, F; Chem Pharm Bull 1986, V34, P4340 HCAPLUS
(2) Bodesheim, U; Pharmazie 1997, V52, P386 HCAPLUS
(3) Eich, E; J Med Chem 1996, V39, P86 HCAPLUS
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L83 ANSWER 10 OF 14 HCAPLUS COPYRIGHT 2004 ACS on STN
     2000:725669 HCAPLUS
AN
ĎΝ
     133:286508
     Entered STN: 13 Oct 2000
ED
     Hydroxymatairesinol preparations in cancer prevention
TI
     Ahotupa, Markku; Eckerman, Christer; Kangas, Lauri;
TN
     Makela, Sari; Saarinen, Niina; Santti, Risto; Warri, Anni
PA
     Hormos Nutraceutical Oy Ltd., Finland
     PCT Int. Appl., 43 pp.
SO
     CODEN: PIXXD2
DT
     Patent
LA
     English
TC
     ICM C07K307-32
     ICS A61K031-00; A23L001-30
     63-6 (Pharmaceuticals)
     Section cross-reference(s): 1, 17
FAN.CNT 2
                                                                  DATE
                                            APPLICATION NO.
     PATENT NO.
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                                            WO 2000-FI181
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                                20001012
PΙ
     WO 2000059946
                         A1
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             CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID,
             IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV,
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MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG,
            SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW,
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                                            US 1999-281094
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                                            EP 2000-909388
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                                20030122
    EP 1165537
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            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO
                                            BR 2000-7187
                                                                   20000309 <--
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    BR 2000007187
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                                20021203
                                            JP 2000-609455
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                         Α
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    ES 2189738
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                                                                   20000309 <--
                                20031120
    AU 767691
                                            NZ 2000-512099
                                                                   20000309 <--
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                                20040130
    NZ 512099
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                                            ZA 2001-4440
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PRAI US 1999-281094
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    WO 2000-FI181
CLASS
                CLASS PATENT FAMILY CLASSIFICATION CODES
 PATENT NO.
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                        C07K307-32
 WO 2000059946
                ICM
                        A61K031-00; A23L001-30
                 ICS
                        A23L001/30B; A61K031/365
 US 6451849
                ECLA
    This invention relates to methods for prevention of cancers, certain
AB
     non-cancer, hormone dependent diseases and/or cardiovascular diseases in a
     person, based on administering of hydroxymatairesinol to said
     person. The invention also concerns a method for increasing the level of
     enterolactone or another metabolite of hydroxymatairesinol
     in a person's serum thereby causing prevention of a cancer or a certain
     non-cancer, hormone dependent disease in a person, based on administering
     of hydroxymatairesinol to said person. Furthermore, this
     invention relates to pharmaceutical prepns., food additives and food
     products comprising hydroxymatairesinol.
     hydroxymatairesinol antitumor hormone disease gynecomastia
ST
IT
    Lignans
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
     (Uses)
        (antioxidant activity of; hydroxymatairesinol prepns. in
        cancer prevention)
TT
     Prostate gland
        (benign hyperplasia; hydroxymatairesinol prepns. in cancer
        prevention)
IT
     Bakery products
        (biscuits; hydroxymatairesinol prepns. in cancer prevention)
IT
     Bakery products
        (cakes; hydroxymatairesinol prepns. in cancer prevention)
IT
     Drug delivery systems
        (carriers; hydroxymatairesinol prepns. in cancer prevention)
IT
     Intestine, neoplasm
     Intestine, neoplasm
        (colon, inhibitors; hydroxymatairesinol prepns. in cancer
        prevention)
     Antitumor agents
IT
        (colon; hydroxymatairesinol prepns. in cancer prevention)
IT
     Cardiovascular system
        (disease; hydroxymatairesinol prepns. in cancer prevention)
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IT
     Hormones, animal, biological studies
     RL: ADV (Adverse effect, including toxicity); BSU (Biological study,
     unclassified); BIOL (Biological study)
        (diseases dependent on; hydroxymatairesinol prepns. in cancer
        prevention)
IT
     Urethra
        (dyssynergia; hydroxymatairesinol prepns. in cancer
        prevention)
IT
     Mammary gland
        (gynecomastia; hydroxymatairesinol prepns. in cancer
        prevention)
IT
     Disease, animal
        (hormone-dependent; hydroxymatairesinol prepns. in cancer
        prevention)
IT
     Antioxidants
     Antitumor agents
     Bread
     Butter
     Candy
     Cardiovascular agents
     Confectionery
     Food
     Food additives
     Margarine
        (hydroxymatairesinol prepns. in cancer prevention)
·IT
     Bladder
        (instability; hydroxymatairesinol prepns. in cancer
        prevention)
IT
     Spruce (Picea abies)
        (lignans of; hydroxymatairesinol prepns. in cancer
        prevention)
IT
     Peroxidation
        (lipid; hydroxymatairesinol prepns. in cancer prevention)
TT
     Lipoproteins
     RL: ADV (Adverse effect, including toxicity); FMU (Formation,
     unclassified); BIOL (Biological study); FORM (Formation, nonpreparative)
        (low-d., oxidation products; hydroxymatairesinol prepns. in
        cancer prevention)
IT
     Urinary tract
        (lower, disease; hydroxymatairesinol prepns. in cancer
        prevention)
IT
     Antitumor agents
        (mammary gland; hydroxymatairesinol prepns. in cancer
        prevention)
IT
     Breakfast cereal
        (muesli; hydroxymatairesinol prepns. in cancer prevention)
IT
     Mammary gland
     Mammary gland
     Prostate gland
     Prostate gland
        (neoplasm, inhibitors; hydroxymatairesinol prepns. in cancer
        prevention)
IT
     Bladder
        (obstruction; hydroxymatairesinol prepns. in cancer
        prevention)
IT
     Blood serum
        (oxidized LDL of; hydroxymatairesinol prepns. in cancer
        prevention)
IT
     Pigments, nonbiological
        (oxidation of; hydroxymatairesinol prepns. in cancer prevention)
IT
     Vitamins
     RL: RCT (Reactant); RACT (Reactant or reagent)
        (oxidation of; hydroxymatairesinol prepns. in cancer prevention)
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huynh - 09 / 991971 IT Lipids, reactions RL: RCT (Reactant); RACT (Reactant or reagent) (peroxidn.; hydroxymatairesinol prepns. in cancer prevention) Antitumor agents TT(prostate gland; hydroxymatairesinol prepns. in cancer prevention) TΤ Milk preparations (yogurt; hydroxymatairesinol prepns. in cancer prevention) 20268-71-7, Hydroxymatairesinol IT RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses) (hydroxymatairesinol prepns. in cancer prevention) IT78473-71-9, Enterolactone RL: BOC (Biological occurrence); BSU (Biological study, unclassified); MFM

(Metabolic formation); THU (Therapeutic use); BIOL (Biological study); FORM (Formation, nonpreparative); OCCU (Occurrence); USES (Uses) (hydroxymatairesinol prepns. in cancer prevention)

9039-48-9, Aromatase IT

RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)

(inhibitors; hydroxymatairesinol prepns. in cancer prevention)

THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD RE.CNT RE

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- 20268-71-7, Hydroxymatairesinol RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)

(hydroxymatairesinol prepns. in cancer prevention)

RN20268-71-7 HCAPLUS

2(3H)-Furanone, dihydro-4-[(S)-hydroxy(4-hydroxy-3-methoxyphenyl)methyl]-3-CN[(4-hydroxy-3-methoxyphenyl)methyl]-, (3R,4R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

78473-71-9, Enterolactone IT

RL: BOC (Biological occurrence); BSU (Biological study, unclassified); MFM (Metabolic formation); THU (Therapeutic use); BIOL (Biological study); FORM (Formation, nonpreparative); OCCU (Occurrence); USES (Uses)

(hydroxymatairesinol prepns. in cancer prevention)

RN78473-71-9 HCAPLUS

2(3H)-Furanone, dihydro-3,4-bis[(3-hydroxyphenyl)methyl]-, (3R,4R)-rel-CN (CA INDEX NAME)

Relative stereochemistry.

L83 ANSWER 11 OF 14 HCAPLUS COPYRIGHT 2004 ACS on STN

AN 2000:517175 HCAPLUS

DN 133:344260

ED Entered STN: 31 Jul 2000

TI Hydroxymatairesinol, a novel enterolactone precursor with antitumor properties from a coniferous tree (Picea abies)

AU Saarinen, N. M.; Warri, A.; Makela, S. I.; Eckerman, C.; Reunanen, M.; Ahotupa, M.; Salmi, S. M.; Franke, A. A.; Kangas, L.; Santti, R.

CS Department of Anatomy and Medical Research Laboratory, University of Turku, Turku, FIN-20520, Finland

SO Nutrition and Cancer (2000), 36(2), 207-214 CODEN: NUCADQ; ISSN: 0163-5581

PB Lawrence Erlbaum Associates, Inc.

DT Journal

LA English

CC 1-6 (Pharmacology)

Section cross-reference(s): 11

The plant lignan hydroxymatairesinol (HMR) was extracted from Norway AB spruce (P. abies) and its metabolism and biol. actions were studied in animals. HMR, the most abundant single component of spruce lignans, was metabolized to enterolactone (ENL) as the major metabolite in rats after oral administration. The amts. of urinary ENL increased with the dose of HMR (3-50 mg/kg), and only minor amts. of unmetabolized HMR isomers and other lignans were found in urine. HMR (15 mg/kg/day for 51 days, orally) decreased the number of growing tumors and increased the proportion of regressing and stabilized tumors in the rat dimethylbenz[a]anthracene-induced mammary tumor model. HMR (50 mg/kg) had no estrogenic or antiestrogenic activity in the uterine growth test in immature rats. HMR also produced no antiandrogenic responses in the growth of accessory sex glands in adult male rats. Neither ENL nor enterodiol had estrogenic or antiestrogenic activity via the classical α - or β -type estrogen receptor-mediated pathway in vitro at <1.0 μ M. HMR was an effective antioxidant in vitro.

ST hydroxymatairesinol enterolactone antitumor antioxidant Picea abies

IT Androgens

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(antiandrogens; antitumor, antioxidant, and other properties of hydroxymatairesinol, a novel enterolactone precursor, from Picea abies)

IT Estrogens

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(antiestrogens; antitumor, antioxidant, and other properties of hydroxymatairesinol, a novel enterolactone precursor, from Picea abies)

IT Antioxidants

Antitumor agents

```
Spruce (Picea abies)
        (antitumor, antioxidant, and other properties of
        hydroxymatairesinol, a novel enterolactone precursor,
        from Picea abies)
IT
     RL: BAC (Biological activity or effector, except adverse); BOC (Biological
     occurrence); BSU (Biological study, unclassified); PRP (Properties); PUR
     (Purification or recovery); THU (Therapeutic use); BIOL (Biological
     study); OCCU (Occurrence); PREP (Preparation); USES (Uses)
        (antitumor, antioxidant, and other properties of
        hydroxymatairesinol, a novel enterolactone precursor,
        from Picea abies)
IT
     Estrogens
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
        (antitumor, antioxidant, and other properties of
        hydroxymatairesinol, a novel enterolactone precursor,
        from Picea abies)
     Estrogen receptors
IT
     RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL
     (Biological study); PROC (Process)
        (hydroxymatairesinol from Picea abies effect on)
     80226-00-2, Enterodiol
IT
     RL: BAC (Biological activity or effector, except adverse); BPR (Biological
     process); BSU (Biological study, unclassified); MFM (Metabolic formation);
     BIOL (Biological study); FORM (Formation, nonpreparative); PROC (Process)
        (antitumor, antioxidant, and other properties of
        hydroxymatairesinol and its metabolite enterodiol, from Picea
        abies)
     78473-71-9, Enterolactone
TT
     RL: BAC (Biological activity or effector, except adverse); BPR (Biological
     process); BSU (Biological study, unclassified); MFM (Metabolic formation);
     BIOL (Biological study); FORM (Formation, nonpreparative); PROC (Process)
        (antitumor, antioxidant, and other properties of
        hydroxymatairesinol and its metabolite enterolactone,
        from Picea abies)
     20268-71-7P, Hydroxymatairesinol
IT
     RL: BAC (Biological activity or effector, except adverse); BOC (Biological
     occurrence); BSU (Biological study, unclassified); PRP (Properties); PUR
     (Purification or recovery); THU (Therapeutic use); BIOL
     (Biological study); OCCU (Occurrence); PREP (Preparation); USES (Uses)
         (antitumor, antioxidant, and other properties of
        hydroxymatairesinol, a novel enterolactone precursor,
        from Picea abies)
     78473-71-9, Enterolactone
TT
     RL: BAC (Biological activity or effector, except adverse); BPR (Biological
     process); BSU (Biological study, unclassified); MFM (Metabolic formation);
     BIOL (Biological study); FORM (Formation, nonpreparative); PROC (Process)
         (antitumor, antioxidant, and other properties of
        hydroxymatairesinol and its metabolite enterolactone,
        from Picea abies)
     78473-71-9 HCAPLUS
RN
     2(3H)-Furanone, dihydro-3,4-bis[(3-hydroxyphenyl)methyl]-, (3R,4R)-rel-
CN
     (9CI) (CA INDEX NAME)
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Relative stereochemistry.

IT 20268-71-7P, Hydroxymatairesinol

RL: BAC (Biological activity or effector, except adverse); BOC (Biological occurrence); BSU (Biological study, unclassified); PRP (Properties); PUR (Purification or recovery); THU (Therapeutic use); BIOL (Biological study); OCCU (Occurrence); PREP (Preparation); USES (Uses)

(antitumor, antioxidant, and other properties of

hydroxymatairesinol, a novel enterolactone precursor,

from Picea abies)

RN 20268-71-7 HCAPLUS

CN 2(3H)-Furanone, dihydro-4-[(S)-hydroxy(4-hydroxy-3-methoxyphenyl)methyl]-3-[(4-hydroxy-3-methoxyphenyl)methyl]-, (3R,4R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

L83 ANSWER 12 OF 14 HCAPLUS COPYRIGHT 2004 ACS on STN

AN 1992:649878 HCAPLUS

DN 117:249878

ED Entered STN: 26 Dec 1992

TI Effect of mammalian lignans of fMLP-induced oxidative bursts in human polymorphonuclear leukocytes

AU Morikawa, Masako; Fukuchi, Kazunori; Inoue, Michiko; Tsuboi, Minoru

CS (Dep. Pharmacol., Tokyo Coll. Pharm., Hachioji, 192-03, Japan

SO Journal of Pharmacy and Pharmacology (1992), 44(10), 859-61

CODEN: JPPMAB; ISSN: 0022-3573

DT Journal

LA English

CC 15-10 (Immunochemistry)

AB The effects of the mammalian lignans, enterolactone, prestegane B and 2,3-dibenzylbutane-1,4-diol (DBB) were examined on superoxide production and luminol-dependent chemiluminescence (LCL) response in human polymorphonuclear leukocytes (PMNs). The three lignans had no direct effect on the responses of human PMNs. DBB and prestegane B enhanced the superoxide production and LCL response induced by formylmethionyl-leucylphenyl-alanine (fMLP), but enterolactone inhibited fMLP-induced effects. The effects of DBB were stronger than those of prestegane B and the effects of DBB were inhibited by bromophenacyl bromide, mepacrine, N-(6-aminophenyl)-5-chloro-1-naphthalene sulfonamide and trifluoroperazine, but not by gossypol, nordihydroguaretic acid, indomethacin, staurosporine, 1-(5-isoquinolinesulfphonyl)-2methylpiperazine dihydrochloride or (R,S)-2-methoxy-3-(octadecylcarbamoyloxy)-propyl-2-(2-thiazolio)-ethylphosphate. These results suggest that DBB primes the responses of human PMNs, and the priming effect is caused by the activation of phospholipase A2- and

Ca2+-calmodulin-pathways, but not by the activation of lipoxygenase, cyclo-oxygenase and protein kinase C or by the release of platelet activating factor.

ST mammal lignan formyl peptide neutrophil respiration

IT Animal respiration

(burst, formyl peptide-induced, of human polymorphonuclear leukocytes, lignans effect on)

IT Hypohalous acids

RL: FORM (Formation, nonpreparative)

(formation of, formyl peptide-induced, in human polymorphonuclear leukocyte respiratory burst, lignans effect on)

IT Lignans

RL: BIOL (Biological study)

(formyl peptide-induced oxidative burst by human polymorphonuclear leukocytes response to)

IT Calmodulins

RL: BIOL (Biological study)

(in lignan modulation of formyl peptide-induced human polymorphonuclear leukocyte oxidative burst)

IT Neutrophil

(respiratory burst of human, formyl peptide-induced, lignans effect on)

IT Leukocyte

(polymorphonuclear, respiratory burst of human, formyl peptide-induced, lignans effect on)

IT 11062-77-4, Superoxide

RL: FORM (Formation, nonpreparative)

(formation of, formyl peptide-induced, in human polymorphonuclear leukocyte respiratory burst, lignans effect on)

IT 78473-71-9, Enterolactone 93376-04-6, Prestegane B

101787-58-0, 2,3-Dibenzylbutane-1,4-diol

RL: BIOL (Biological study)

(formyl peptide-induced oxidative burst by human polymorphonuclear leukocytes response to)

IT 9001-84-7, Phospholipase A2

RL: BIOL (Biological study)

(in lignan modulation of formyl peptide-induced human polymorphonuclear leukocyte oxidative burst)

IT 59880-97-6

RL: BIOL (Biological study)

(oxidative burst induction by, in human polymorphonuclear leukocytes, lignans effect on)

IT 78473-71-9, Enterolactone

RL: BIOL (Biological study)

(formyl peptide-induced oxidative burst by human polymorphonuclear leukocytes response to)

RN 78473-71-9 HCAPLUS

CN 2(3H)-Furanone, dihydro-3,4-bis[(3-hydroxyphenyl)methyl]-, (3R,4R)-rel-(9CI) (CA INDEX NAME)

Relative stereochemistry.

L83 ANSWER 13 OF 14 HCAPLUS COPYRIGHT 2004 ACS on STN

AN 1991:23554 HCAPLUS

DN 114:23554

ED Entered STN: 26 Jan 1991

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TI Preparation of dibenzylbutanediol and dibenzyltetrahydrofuran derivatives as immunosuppressants
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IN Oka, Kitaro; Hirano, Toshihiko; Naito, Takashi; Hosaka, Kunio

PA Tsumura and Co., Japan

SO Jpn. Kokai Tokkyo Koho, 21 pp.

CODEN: JKXXAF

DT Patent

LA Japanese IC ICM A61K031-05

ICS A61K031-075; A61K031-22; A61K031-34

ICA C07C033-24; C07C039-15; C07C043-20; C07C069-21; C07D307-10; C07D307-33

CC 25-18 (Benzene, Its Derivatives, and Condensed Benzenoid Compounds) Section cross-reference(s): 1, 27, 63

FAN.CNT 1

PATENT NO.		KIND	DATE	APPLICATION NO.	DATE
PI JP 02040323		A2	19900209	JP 1988-186853	19880728
PRAI JP 1988-186853			19880728		
CLASS				•	
PATENT NO.	CLASS	PATENT	FAMILY CLASS	IFICATION CODES	
JP 02040323	A61K031-05				
ICS A61K031-075; A61K031-22; A61					
	ICA	C07C033	-24: C07C039	-15: C07C043-20:	C07C069-21:

C07D307-10; C07D307-33

OS MARPAT 114:23554

GΙ

$$B1$$
 CH_2
 CH_2
 $B1$
 $B1$

The title compds. (I, II; A, A1, A2 = H, OH, MeO; A3 = H, Me, Ac; B1 = H, MeO; Z = O, 2H) were prepared and formulated as immunosuppressants. A solution of hydrocinnamic acid in THF was added to BuLi-hexane at -72° with stirring under Ar, the solution warmed to -10°, cooled to -62°, a solution of iodine in THF was added to give (±)-2,3-dibenzylsuccinic acid, which was esterified with MeI in DMF under Ar to give the di-Me ester (III). Reduction of III gave diol (±)-I (A = A1 = A2 = A3 = H), which inhibited mitogen-stimulated human peripheral lymphocyte proliferation by 56.8%. Also prepared and tested were 17 addnl. I and II. Tablet, granular, and injection formulations were also given.

II

- ST immunosuppressant dibenzylbutanediol dibenzyltetrahydrofuran prepn; benzylbutanediol prepn immunosuppressant; benzyltetrahydrofuran prepn immunosuppressant
- IT Immunosuppressants

(dibenzylbutanediol and dibenzyltetrahydrofuran derivs.)

IT 501-52-0, Hydrocinnamic acid

RL: RCT (Reactant); RACT (Reactant or reagent)
 (coupling reaction of)

IT 2316-26-9, 3,4-Dimethoxycinnamic acid 6099-04-3, m-Methoxycinnamic acid

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RL: RCT (Reactant); RACT (Reactant or reagent)
        (hydrogenation of, in preparation of immunosuppressants)
     2107-70-2P, 3,4-Dimethoxyhydrocinnamic acid 10516-71-9P,
TΤ
     3-Methoxydihydrocinnamic acid
     RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT
     (Reactant or reagent)
        (preparation and coupling reaction of, in preparation of immunosuppressants)
     93609-04-2P
IT
     RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT
     (Reactant or reagent)
        (preparation and dehydration of, in preparation of immunosuppressants)
                   93578-39-3P 119516-58-4P
                                               126965-29-5P
                                                               126965-30-8P
IT
     126965-33-1P
     RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT
     (Reactant or reagent)
        (preparation and esterification of, in preparation of immunosuppressants)
IT
     121955-01-9P
     RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT
     (Reactant or reagent)
        (preparation and etherification of, in preparation of immunosuppressants)
                   126981-89-3P
     126965-31-9P
IT
     RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT
     (Reactant or reagent)
        (preparation and lactonization of, in preparation of immunosuppressants)
                   119516-59-5P 121955-10-0P 126965-28-4P
                                                               126965-34-2P
     81436-89-7P
TΤ
     RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT
     (Reactant or reagent)
        (preparation and reduction of, in preparation of immunosuppressants)
                                 78473-70-8P 78473-71-9P
                   77756-23-1P
IT
     77756-22-0P
                                                 121955-04-2P
                                                                 121955-05-3P
     93451-90-2P
                   119516-60-8P
                                  121851-41-0P
                                   121955-09-7P
                                                  121986-75-2P
                                                                 122045-61-8P
     121955-06-4P
                    121955-07-5P
     122045-63-0P
                    123808-59-3P
                                   123877-50-9P
                                                  131049-50-8P
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); SPN (Synthetic preparation); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (preparation of, as immunosuppressant)
IT
     78473-71-9P
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); SPN (Synthetic preparation); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (preparation of, as immunosuppressant)
     78473-71-9 HCAPLUS
RN
     2(3H)-Furanone, dihydro-3,4-bis[(3-hydroxyphenyl)methyl]-, (3R,4R)-rel-
CN
           (CA INDEX NAME)
```

Relative stereochemistry.

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L83 ANSWER 14 OF 14 HCAPLUS COPYRIGHT 2004 ACS on STN

AN 1982:17711 HCAPLUS

DN 96:17711

ED Entered STN: 12 May 1984

TI Lignan formation in man. Microbial involvement and possible roles in relation to cancer

AU Setchell, K. D. R.; Borriello, S. P.; Gordon, H.; Lawson, A. M.; Harkness, R.; Morgan, D. M. L.; Kirk, D. N.; Adlercreutz, H.; Anderson, L. C.;
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Axelson, M.
     Div. Clin. Chem., Clin. Res. Cent., Harrow, HA1 3UJ, UK
CS
     Lancet (1981), 2(8236), 4-7
SO
     CODEN: LANCAO; ISSN: 0023-7507
DT
     Journal
     English
LA
CC
     13-2 (Mammalian Biochemistry)
     Section cross-reference(s): 2, 4, 10
     Studies of the formation of 2 lignans (named enterolactone and
AB
     enterodiol) in man, by means of selective antibiotic administration,
     confirmed that these new compds. are formed by intestinal microflora.
     Bacteriol. studies of stools collected after metronidazole administration
     indicated that clostridia may be responsible for the formation of these
     highly aromatic compds. The role of lignans in intestinal cancer is
     discussed.
ST
     lignan formation intestine microorganism cancer; Clostridium intestine
     lignan formation cancer; urine feces enterolactone enterodiol
     Nomenclature, new natural products
IT
        (enterodiol)
IT
     Feces
     Urine
        (enterodiol and enterolactone of, intestinal microorganisms
        in relation to)
IT
     Nomenclature, new natural products
        (enterolactone)
IT
     Lymphocyte
        (enterolactone toxicity to)
IT
     Lignans
     RL: FORM (Formation, nonpreparative)
        (formation of, intestinal microflora in, cancer in relation to)
IT
     Clostridium
        (intestinal, in lignan formation, cancer in relation to)
IT
     Neoplasm
        (of intestine, lignans in relation to)
IT
     Intestine, neoplasm
        (cancer, lignans in relation to)
IT
     Microorganism
        (intestinal, in lignan formation, cancer in relation to)
IT
     78473-71-9
                80226-00-2
     RL: FORM (Formation, nonpreparative)
        (formation of, intestinal microorganisms in relation to)
IT
     78473-71-9
     RL: FORM (Formation, nonpreparative)
        (formation of, intestinal microorganisms in relation to)
RN
     78473-71-9 HCAPLUS
     2(3H)-Furanone, dihydro-3,4-bis[(3-hydroxyphenyl)methyl]-, (3R,4R)-rel-
CN
           (CA INDEX NAME)
     (9CI)
```

Relative stereochemistry.

=> => d all hitstr tot 188

L88 ANSWER 1 OF 2 HCAPLUS COPYRIGHT 2004 ACS on STN AN 1997:22530 HCAPLUS

```
126:139494
DN
     Entered STN: 15 Jan 1997
ED
     Studies on differentiation inducers. VI. Lignan derivatives from Arctium
TT
     Umehara, Kaoru; Nakamura, Mitsuhiro; Miyase, Toshio; Kuroyanagi, Masanori;
ΑU
     Uneo, Akira
     Sch. Pharm. Sci., Univ. Shizuoka, Shizuoka, 422, Japan
CS
     Chemical & Pharmaceutical Bulletin (1996), 44(12), 2300-2304
SO
     CODEN: CPBTAL; ISSN: 0009-2363
PB
     Pharmaceutical Society of Japan
DT
     Journal
LA
     English
CC
     1-3 (Pharmacology)
     Section cross-reference(s): 11, 26
     In the previous paper, we reported the differentiation inducing activities
AB
     of lignoids from Arctium Fructus (the fruits of Arctium lappa L.,
     compositae) against mouse myeloid leukemia cells (M1). We reinvestigated
     the active components of this extract and isolated three new dilignans.
     Furthermore, structure modifications were carried out using the most
     active lignan (arctigenin) and its structure-activity relationship was
     investigated. Its aliphatic esters were more effective in inducing the
     differentiation of M1 cells than its aromatic esters. Especially n-docanoate,
     which was the most active derivative, induced more than half of the M1 cells
     into phagocytic cells at a concentration of 2 \mu M.
     Arctium lignan deriv prepn differentiation inducer; leukemia cell
ST
     differentiation inducer lignan deriv
     New natural products
IT
        (arctignan F (lignan))
     New natural products
IT
        (arctignan G (lignan))
     New natural products
TT
        (arctignan H (lignan))
     Cell differentiation
TT
        (inducers; lignans from Arctium Fructus as myeloid leukemia
        differentiation inducers)
     Antitumor agents
IT
        (leukemia; lignans from Arctium Fructus as myeloid leukemia
        differentiation inducers)
IT
     Arctium
     Macrophage
       Phagocytosis
     Structure-activity relationship
        (lignans from Arctium Fructus as myeloid leukemia differentiation
TT
     Lignans
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); BIOL (Biological study)
       (lignans from Arctium Fructus as myeloid leukemia differentiation
        inducers)
IT
     Leukemia
        (myelogenous; lignans from Arctium Fructus as myeloid leukemia
        differentiation inducers)
                               62333-08-8P, Lappaol A
     580-72-3P, Matairesinol
                              64855-00-1P, Lappaol C
                                                        131400-96-9P, Isolappaol
     62359-60-8P, Lappaol B
                                     186541-89-9P, Arctignan G
                                                                 186543-11-3P,
         186541-65-1P, Arctignan F
     Arctignan H
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); PUR (Purification or recovery); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (lignans from Arctium Fructus as myeloid leukemia differentiation
        inducers)
     69232-85-5P
                   74861-36-2P
IT
```

RL: BAC (Biological activity or effector, except adverse); BSU (Biological

study, unclassified); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)

(lignans from Arctium Fructus as myeloid leukemia differentiation inducers)

IT 7770-78-7 20362-31-6

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); RCT (Reactant); THU (Therapeutic use); BIOL (Biological study); RACT (Reactant or reagent); USES (Uses)

(lignans from Arctium Fructus as myeloid leukemia differentiation inducers)

IT 25488-59-9P 73354-08-2P 119069-38-4P 186449-80-9P 186449-81-0P 186449-82-1P 186449-83-2P 186449-84-3P 186449-85-4P 186449-86-5P 186449-89-8P 186449-90-1P 186449-87-6P 186449-88-7P 186449-91-2P 186449-92-3P 186449-93-4P 186449-94-5P 186449-95-6P 186449-96-7P 186583-53-9P 186449-97-8P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(lignans from Arctium Fructus as myeloid leukemia differentiation inducers)

IT 41328-76-1 69394-17-8, Lappaol F

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(lignans from Arctium Fructus as myeloid leukemia differentiation inducers)

RE.CNT 13 THERE ARE 13 CITED REFERENCES AVAILABLE FOR THIS RECORD RE

(1) Freudenberg, K; Chem Ber 1957, V90, P2857

- (2) Hartwell, J; Fortschr Chem Org Naturst 1958, V15, P83 HCAPLUS
- (3) Kupchan, S; J Am Chem Soc 1973, V95, P1335 HCAPLUS
- (4) Landais, Y; Tetrahedron 1991, V47, P3787 HCAPLUS
- (5) Magae, J; Agric Biol Chem 1988, V52, P3143 HCAPLUS
- (6) Miyamura, C; FEBS Lett 1988, V234, P17
- (7) Nishibe, S; Chem Pharm Bull 1981, V29, P2078 HCAPLUS
- (8) Sugiyama, S; Chem Pharm Bull 1993, V41, P714 HCAPLUS
- (9) Umehara, K; Chem Pharm Bull 1992, V40, P401 HCAPLUS
- (10) Umehara, K; Chem Pharm Bull 1993, V41, P1774 HCAPLUS
- (11) Umehara, K; Chem Pharm Bull 1994, V42, P611 HCAPLUS
- (12) Umehara, K; Chem Pharm Bull 1995, V43, P1565 HCAPLUS
- (13) Yamamoto, Y; Exp Cell Res 1986, V164, P97
- IT 580-72-3P, Matairesinol

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); PUR (Purification or recovery); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(lignans from Arctium Fructus as myeloid leukemia differentiation inducers)

RN 580-72-3 HCAPLUS

CN 2(3H)-Furanone, dihydro-3,4-bis[(4-hydroxy-3-methoxyphenyl)methyl]-, (3R,4R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

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ANSWER 2 OF 2 HCAPLUS COPYRIGHT 2004 ACS on STN
    1994:431077 HCAPLUS
AN
     121:31077
DN
     Entered STN: 23 Jul 1994
ED
     Studies on differentiation-inducers from Arctium Fructus
TI
     Umehara, Kaoru; Sugawa, Ariko; Kuroyanagi, Masanori; Ueno, Akira; Taki,
ΑU
     Sch. Pharm. Sci., Univ. Shizuoka, Shizuoka, 422, Japan
     Chemical & Pharmaceutical Bulletin (1993), 41(10), 1774-9
SO.
     CODEN: CPBTAL; ISSN: 0009-2363
T.A
     Journal
     English
LA
     11-1 (Plant Biochemistry)
CC
     Section cross-reference(s): 1, 26
     In the course of studying differentiation-inducers from plants, their
ΑB
     isolation was performed from the methanolic extract of Arctium Fructus (the
     fruits of Arctium lappa L., Compositae), and then their phagocytic
     activity on differentiated mouse myeloid leukemia cells (M1) was
     monitored. Thirteen compds., including five new ones (arctignans A-E),
     were isolated as differentiation-inducers toward M1 cells. These
     consisted of two lignans, eight sesquilignans and three dilignans.
     Arctigenin (2) was the most effective compound of all those isolated, and it
     induced differentiation of M1 cells at a concentration 0.5 \mu M. Sesquilignans
     were less effective than lignans and dilignans showed even weaker
     activity. These lignoids were inactive towards a human acute
     promyelocytic leukemia cell line (HL-60).
     Arctium arctignan neoplasm inhibitor cell differentiation
ST
     Nomenclature, new natural products
IT
        (arctignan A (lignan))
     Nomenclature, new natural products
IT
        (arctignan B (lignan))
     Nomenclature, new natural products
IT
         (arctignan C (lignan))
     Nomenclature, new natural products
IT
         (arctignan D (lignan))
     Nomenclature, new natural products
IT
         (arctignan E (lignan))
     Lignans
IT
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); BIOL (Biological study)
         (from Arctium lappa, isolation and phagocytic activity of, on
        differentiated myeloid leukemia cells)
     Cell differentiation
IT
         (inducers, lignans from Arctium lappa as, of tumor cells)
     Molecular structure, natural product
IT
         (of arctignan A (lignan))
     Molecular structure, natural product
IT
         (of arctignan B (lignan))
     Molecular structure, natural product
IT
         (of arctignan C (lignan))
     Molecular structure, natural product
IT
         (of arctignan D (lignan))
     Molecular structure, natural product
IT
         (of arctignan E (lignan))
     Lignans
IT
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); BIOL (Biological study)
         (di-, from Arctium lappa, isolation and phagocytic activity
         of, on differentiated myeloid leukemia cells)
     Neoplasm inhibitors
IT
         (leukemia, lignans from Arctium lappa as)
IT
     Lignans
```

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study)

(sesqui-, from Arctium lappa, isolation and **phagocytic** activity of, on differentiated myeloid leukemia cells)

IT 580-72-3 7770-78-7, Arctigenin 20362-31-6 62333-08-8 62359-60-8 64855-00-1 64855-02-3 69394-17-8 131400-96-9 155661-08-8, Arctignan A 155661-09-9, Arctignan B 155661-10-2,

Arctignan C 155661-11-3, Arctignan D 155661-12-4, Arctignan E

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study)

(from Arctium lappa, isolation and **phagocytic** activity of, on differentiated myeloid leukemia cells)

25488-59-9P 69232-85-5P 74861-36-2P 119069-38-4P

RL: SPN (Synthetic preparation); PREP (Preparation)

(preparation and phagocytic activity of, on differentiated myeloid leukemia cells)

IT 580-72-3

IT

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study)

(from Arctium lappa, isolation and **phagocytic** activity of, on differentiated myeloid leukemia cells)

RN 580-72-3 HCAPLUS

CN 2(3H)-Furanone, dihydro-3,4-bis[(4-hydroxy-3-methoxyphenyl)methyl]-, (3R,4R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

=> => fil medline

FILE 'MEDLINE' ENTERED AT 14:32:05 ON 30 SEP 2004

FILE LAST UPDATED: 29 SEP 2004 (20040929/UP). FILE COVERS 1951 TO DATE.

On February 29, 2004, the 2004 MeSH terms were loaded. See HELP RLOAD for details. OLDMEDLINE now back to 1951.

MEDLINE thesauri in the /CN, /CT, and /MN fields incorporate the MeSH 2004 vocabulary. See http://www.nlm.nih.gov/mesh/ and http://www.nlm.nih.gov/pubs/techbull/nd03/nd03_mesh.html for a description of changes.

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> d all tot 1108

L108 ANSWER 1 OF 5 MEDLINE on STN

AN 2003070677 MEDLINE

DN PubMed ID: 12580104

TI New biflavanones and bioactive compounds from Stellera chamaejasme L.

AU Xu Z H; Qin G W; Li X Y; Xu R S

CS Shanghai Institute of Materia Medica, Shanghai Institutes of Biological Sciences, Chinese Academy of Sciences, Shanghai 200031, China.

```
Yao xue xue bao = Acta pharmaceutica Sinica, (2001 Sep) 36 (9)
SO
     669-71.
    Journal code: 21710340R. ISSN: 0513-4870.
CY
    China
    Journal; Article; (JOURNAL ARTICLE)
DT
     English
LA
FS
     Priority Journals
EM
    200402
    Entered STN: 20030214
ED
    Last Updated on STN: 20040207
    Entered Medline: 20040206
    AIM: To study the chemical constituents of the root of Stellera
AB
    chamaejasme L. METHODS: Various column chromatographies on silica gel and
    RP-18 were employed for isolation and purification. Structures of
     compounds were elucidated by spectral analysis. RESULTS: Eight lignans
    and three biflavonoids possessing a C-3/C-3" linkage were isolated. They
     are ruixianglangdusu A (1) and B (2), 4',4'",5,5",7,7"-hexahydroxy-3,3"-
    biflavone (3), (+)-kusunokinin (4), lirioresinol-B (5), magnolenin C (6),
     (-)-pinoresinol monomethyl ether (7), (-)-pinoresinol (8), (+)-
    matairesinol (9), isohinokinin (10) and (-)-eudesmin (11).
    CONCLUSION: Compounds 1 and 2 are new biflavanones, 1 is enantiomeric to
    known chamaejasmenin C, 4, 6, 8, 9, 10 and 11 were isolated from this
    plant for the first time, and 7 was isolated from natural resources for
     the first time. In vitro bioassays showed that 3 and 8 exhibited
     antibacterial activity, and 1, 2, 9 and 11 exhibited immunomodulatory
     activity.
     Adjuvants, Immunologic: CH, chemistry
CT
     Adjuvants, Immunologic: IP, isolation & purification
      Adjuvants, Immunologic: PD, pharmacology
      Anti-Infective Agents: CH, chemistry
      Anti-Infective Agents: IP, isolation & purification
      Anti-Infective Agents: PD, pharmacology
        B-Lymphocytes: DE, drug effects
      Flavanones: CH, chemistry
     *Flavanones: IP, isolation & purification
      Flavanones: PD, pharmacology
      Molecular Structure
     Plant Roots: CH, chemistry
     *Plants, Medicinal: CH, chemistry
     *Thymelaeaceae: CH, chemistry
     0 (Adjuvants, Immunologic); 0 (Anti-Infective Agents); 0 (Flavanones); 0
CN
     (ruixianglangdusu A); 0 (ruixianglangdusu B)
L108 ANSWER 2 OF 5
                       MEDLINE on STN
     2001451032
                    MEDLINE
AN
DN
     PubMed ID: 11473435
    Anti-AIDS agents. 46. Anti-HIV activity of harman, an anti-HIV principle
TT
     from Symplocos setchuensis, and its derivatives.
     Ishida J; Wang H K; Oyama M; Cosentino M L; Hu C Q; Lee K H
ΑU
    Natural Products Laboratory, School of Pharmacy, University of North
CS
     Carolina, Chapel Hill, North Carolina 27599-7360, USA.
     AI 33066 (NIAID)
NC
     Journal of natural products, (2001 Jul) 64 (7) 958-60.
SO
     Journal code: 7906882. ISSN: 0163-3864.
CY
     United States
     Journal; Article; (JOURNAL ARTICLE)
DT
LA
     English
FS
     Priority Journals
EM
     200110
     Entered STN: 20010813
ED
     Last Updated on STN: 20011015
     Entered Medline: 20011011
    Matairesinol (1) and harman (5), identified from Symplocos
AB
```

setchuensis, were found to inhibit HIV replication in H9 lymphocyte cells. Anti-HIV evaluation of 28 derivatives of 5 revealed that compound 19 showed potent activity with EC(50) and therapeutic index values of 0.037 microM and 210, respectively. CTCheck Tags: Comparative Study; Human; Support, U.S. Gov't, P.H.S. Anti-HIV Agents: CH, chemistry *Anti-HIV Agents: IP, isolation & purification Anti-HIV Agents: PD, pharmacology Chromatography, High Pressure Liquid Drugs, Chinese Herbal: CH, chemistry *Drugs, Chinese Herbal: IP, isolation & purification Drugs, Chinese Herbal: PD, pharmacology Furans: CH, chemistry *Furans: IP, isolation & purification Furans: PD, pharmacology Harmine: AA, analogs & derivatives Harmine: CH, chemistry
*Harmine: IP, isolation & purification Harmine: PD, pharmacology Lignans: CH, chemistry *Lignans: IP, isolation & purification Lignans: PD, pharmacology Lymphocytes: DE, drug effects Lymphocytes: ME, metabolism Molecular Structure *Plants, Medicinal: CH, chemistry Structure-Activity Relationship 442-51-3 (Harmine); 486-84-0 (harman); 580-72-3 (matairesinol) RN0 (Anti-HIV Agents); 0 (Drugs, Chinese Herbal); 0 (Furans); 0 (Lignans); 0 CN (N-butylharman) L108 ANSWER 3 OF 5 MEDLINE on STN 2000296660 MEDLINE AN DNPubMed ID: 10837017 Isoflavonoids and lignans have different potentials to modulate oxidative TIgenetic damage in human colon cells. Pool-Zobel B L; Adlercreutz H; Glei M; Liegibel U M; Sittlingon J; Rowland AU I; Wahala K; Rechkemmer G Department of Nutritional Toxicology, Institute for Nutrition, Friedrich CS Schiller University, Dornburger Strabetae 25, 07743 Jena, Germany... b8pobe@uni-jena.de Carcinogenesis, (2000 Jun) 21 (6) 1247-52. SO Journal code: 8008055. ISSN: 0143-3334. ENGLAND: United Kingdom DT (CLINICAL-TRIAL) Journal; Article; (JOURNAL ARTICLE) (RANDOMIZED CONTROLLED TRIAL) LA English Priority Journals FS 200008 EM ED Entered STN: 20000811 Last Updated on STN: 20000811 Entered Medline: 20000803 Polyphenolic compounds, including isoflavonoids and lignans, have been AB suggested to be chemopreventive on account of antioxidative properties. In this context it is of importance to have knowledge of their ability to reduce oxidative stress within target cells of tumorigenesis. Therefore, we investigated isoflavonoids and lignans for modulation of oxidative genetic damage in mammalian cells. H(2)O(2)-induced damage as well as endogenous DNA strand breaks and oxidized bases were determined after 30 min incubation of human colon cells with polyphenols using various modifications of the microgel electrophoresis assay (Comet assay).

Enterolactone, a mammalian metabolite of plant lignans, was

additionally investigated for modulation of intracellular oxidative stress in NIH 3T3 cells using laser scanning microscopy. In vivo effects of rye crispbread (a source of lignans) were investigated in 12 human volunteers by determining genetic damage in lymphocytes and antioxidant activity in plasma (FRAP assay). Genistein induced DNA breaks in the human tumour cell line HT29 clone 19A (12.5-100 microM). The polyphenols (100 microM) did not reduce damage induced by 150 microM H(2)O(2), indicating that they lacked antioxidative potential. At this concentration enterolactone also had no effect on intracellular oxidative stress induced by 31.25 and 125 microM H(2)O(2). In contrast, enterolactone, dihydrogenistein and formononetin reduced endogenous oxidative DNA damage at 100 microM. Daily ingestion of nine slices (76.5 g/day) of rye crispbread per day (containing 41.8 and 33.0 microg/100 g dry weight secoisolariciresinol and matairesinol, respectively) for 2 weeks did not significantly reduce genetic damage in blood lymphocytes, nor was there a modulation of plasma antioxidant capacity. The moderate effects of high concentrations of the tested compounds on endogenous oxidative DNA damage and failure to prevent H(2)O(2) - induced damage are indicative of only marginal protective potential by antioxidant mechanisms. The genotoxic effects of genistein deserve further investigation. Check Tags: Human; Support, Non-U.S. Gov't 3T3 Cells Animals *Antimutagenic Agents: PD, pharmacology Bread

CT

*Colon: DE, drug effects Colon: PA, pathology Comet Assay

Cross-Over Studies

*DNA Damage

*Flavonoids: PD, pharmacology *Lignans: PD, pharmacology Mice

*Oxidative Stress

Secale cereale

0 (Antimutagenic Agents); 0 (Flavonoids); 0 (Lignans) CN

MEDLINE on STN L108 ANSWER 4 OF 5

MEDLINE AN94366249

PubMed ID: 8084211 DN

Natural flavonoids and lignans are potent cytostatic agents against human ΤI leukemic HL-60 cells.

ΑU Hirano T; Gotoh M; Oka K

Department of Clinical Pharmacology, Tokyo College of Pharmacy, Japan. CS

Life sciences, (1994) 55 (13) 1061-9. SO Journal code: 0375521. ISSN: 0024-3205.

CYENGLAND: United Kingdom

Journal; Article; (JOURNAL ARTICLE) DT

LA English

FS Priority Journals

EΜ 199410

Entered STN: 19941021 EDLast Updated on STN: 19970203 Entered Medline: 19941011

Anti leukemic-cell efficacy of 28 naturally occurring and synthetic flavonoids and 11 naturally occurring lignans on human promyelocytic leukemic cell line HL-60 were examined using MTT assay methods. Differences between anti cell-proliferative activity and cytotoxicity of these compounds were compared with those of 4 clinical anti-cancer agents. Eight of the 28 flavonoids and 4 of the 11 lignans showed considerable suppressive effects on HL-60 cell growth with IC50s ranging from 10-940 ng/ml. Among these compounds, genistein, honokiol, machilin A,

matairesinol, and arctigenin had the strongest effects with IC50s less than 100 ng/ml, which were almost equivalent to the effects of current anti-cancer agents. The flavonoid genistein and the lignans, however, showed little or no cytotoxicity against HL-60 cells as assessed by dye exclusion tests (LC50s > 2,900 ng/ml), whereas the regular anti-cancer agents had potent cytotoxicity. All of the flavonoids and lignans, except for machilin A and arctigenin, were less effective against growth of human T lymphocytic leukemia cell line MOLT-4. In addition, the flavonoid and the lignans showed little or no inhibiting activity on mitogen-induced blastogenesis of human peripheral-blood lymphocytes. lignans and genistein were strongly suppressive against incorporations of [3H] thymidine, [3H] uridine, and [3H] leucine into HL-60 cells. These results showed that some of the naturally occurring flavonoids and lignans inhibited HL-60 cell growth with a non-toxic mechanism, possibly via cessation of DNA, RNA, and/or protein synthesis of the leukemic cells. Check Tags: Comparative Study; Human *Antineoplastic Agents: PD, pharmacology Cell Division: DE, drug effects Drug Screening Assays, Antitumor *Flavonoids: PD, pharmacology Leucine: ME, metabolism *Leukemia, Promyelocytic, Acute: DT, drug therapy Leukemia, Promyelocytic, Acute: ME, metabolism Leukemia, Promyelocytic, Acute: PA, pathology Leukemia, T-Cell: DT, drug therapy Leukemia, T-Cell: PA, pathology *Lignans: PD, pharmacology Lymphocyte Activation: DE, drug effects Lymphocytes: DE, drug effects Lymphocytes: IM, immunology Tetrazolium Salts Thiazoles Thymidine: ME, metabolism Tumor Cells, Cultured: DE, drug effects Uridine: ME, metabolism 298-93-1 (thiazolyl blue); 50-89-5 (Thymidine); 58-96-8 (Uridine); 61-90-5 (Leucine) 0 (Antineoplastic Agents); 0 (Flavonoids); 0 (Lignans); 0 (Tetrazolium Salts); 0 (Thiazoles) MEDLINE on STN L108 ANSWER 5 OF 5 93085549 MEDLINE PubMed ID: 1360514 Effect of mammalian lignans on fMLP-induced oxidative bursts in human polymorphonuclear leucocytes. Morikawa M; Fukuchi K; Inoue M; Tsuboi M Department of Pharmacology, Tokyo College of Pharmacy, Japan. Journal of pharmacy and pharmacology, (1992 Oct) 44 (10) 859-61. Journal code: 0376363. ISSN: 0022-3573. ENGLAND: United Kingdom Journal; Article; (JOURNAL ARTICLE) English Priority Journals 199301 Entered STN: 19930129 Last Updated on STN: 19950206 Entered Medline: 19930104 We examined the effects of mammalian lignans, enterolactone, prestegane B and 2,3-dibenzylbutane-1,4-diol (DBB) on superoxide production and luminol-dependent chemiluminescence (LCL) response in human polymorphonuclear leucocytes (PMNs). The three lignans had no direct effect on the responses of human PMNs. DBB and prestegane B enhanced the

superoxide production and LCL response induced by formylmethionyl-leucyl-

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LA

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AB

phenylalanine (fMLP), but enterolactone inhibited fMLP-induced effects. The effects of DBB were stronger than those of prestegane B and the effects of DBB were inhibited by bromophenacyl bromide, mepacrine, N-(6-aminophenyl)-5-chloro-1-naphthalene, sulphonamide and trifluoroperazine, but not by gossypol, nordihydroguaretic acid, indomethacin, staurosporine, 1-(5-isoquinolinesulphonyl)-2methylpiperazine dihydrochloride or (R,S)-2-methoxy-3-(octadecylcarbamoyloxy)-propyl-2-(2-thiazoli o)-ethylphosphate. These results suggest that DBB primes the responses of human PMNs, and the priming effect is caused by the activation of phospholipase A2--and Ca(2+)-calmodulin-pathways, but not by the activation of lipoxygenase, cyclo-oxygenase and protein kinase C or by the release of platelet activating factor. Check Tags: Human; In Vitro Chemiluminescence Lignans *Lignin: PD, pharmacology *N-Formylmethionine Leucyl-Phenylalanine: PD, pharmacology *Neutrophils: DE, drug effects *Respiratory Burst: DE, drug effects Superoxides: AN, analysis 11062-77-4 (Superoxides); 59880-97-6 (N-Formylmethionine Leucyl-Phenylalanine); 9005-53-2 (Lignin) 0 (Lignans) => => fil bios is 'BIOS' IS AN AMBIGUOUS FILE OR CLUSTER NAME - Bioscience Literature Cluster BIOSCIENCE - The BIOSIS Previews(R)/RN File 1969-present BIOSIS ENTER FILE OR CLUSTER NAME (IGNORE):end => fil biosis FILE 'BIOSIS' ENTERED AT 14:34:56 ON 30 SEP 2004 Copyright (c) 2004 The Thomson Corporation. FILE COVERS 1969 TO DATE. CAS REGISTRY NUMBERS AND CHEMICAL NAMES (CNs) PRESENT FROM JANUARY 1969 TO DATE. RECORDS LAST ADDED: 29 September 2004 (20040929/ED) FILE RELOADED: 19 October 2003. => d all tot 1116 L116 ANSWER 1 OF 7 BIOSIS COPYRIGHT (c) 2004 The Thomson Corporation. on STN 2004:386345 BIOSIS PREV200400386057 Prenatal developmental toxicity study with 7-hydroxymatairesinol potassium acetate (HMRlignan) in rats. Wolterbeek, A. P. M. [Reprint Author]; Roberts, A.; Korte, H.; Unkila, M.; Waalkens-Berendsen, D. H. Toxicol and Appl Pharmacol Dept, TNO Nutr and Food Res, Zeist, Netherlands wolterbeek@voeding.tno.nl Regulatory Toxicology and Pharmacology, (August 2004) Vol. 40, No. 1, pp. 1-8. print. CODEN: RTOPDW. ISSN: 0273-2300. Article English Entered STN: 29 Sep 2004 Last Updated on STN: 29 Sep 2004 Plant lignan 7-hydromatairesinol, a novel precursor of the mammalian

lignan enterolactone was evaluated in a prenatal developmental toxicity study conducted in the Wistar, rat. Mated female rats were fed diets containing 0, 0.25, 1, and 4% (w/w) of 7-hydroxymatairesinol in the form of potassium acetate complex (HMR lignan; potassium acetate level approximately 20% w/w within the preparation) for days 0-21 of gestation. Test substance intake was calculated to be 0.14-0.18, 0.46-0.74, and 1.19-2.93 g/kg body weight/day for the low, mid, and high-dose groups, respectively. The rats were sacrificed on day 21 of the gestation period and examined for standard parameters of reproductive performance (fecundity index, gestation index, number of corpora lutea, number of implantations, pre- and post-implantation loss, number of earlyand late resorptions, number of live- and dead fetuses, sex-ration and the weight of the reproductive organs). The fetuses were examined for external, visceral, and skeletal alterations. The results from this study showed no effects on reproductive performance or any treatment related findings following external, visceral, and skeletal examination of the fetuses. However, approximately half of the mated dams of the high-dose failed to thrive due to an unexpected large decrease in their food intake, and were sacrificed early. Body weights of the remaining animals of the high-dose group were decreased. Food consumption was decreased in all treatment groups during the first three days of the gestation period as a result of decreased palatability of the feed. In conclusion, the no-observed-effect level (NOEL) for maternal effects was 1%, whereas the NOEL for fetal development following daily oral HMR lignan administration throughout the gestation was equivalent to 4% in the diet. Copyright 2004 Elsevier Inc. All rights reserved.

CC Nutrition - General studies, nutritional status and methods 13202
Reproductive system - Physiology and biochemistry 16504
Toxicology - General and methods 22501
Development and Embryology - General and descriptive 25502

IT Major Concepts

Nutrition; Reproduction; Toxicology

IT Chemicals & Biochemicals

7-hydroxymatairesinol potassium acetate; lignan enterolactone; plant lignan; potassium acetate

IT Miscellaneous Descriptors

prenatal developmental toxicity study; reproductive performance ORGN Classifier

Muridae 86375

Super Taxa

Rodentia; Mammalia; Vertebrata; Chordata; Animalia

Organism Name

Wistar rat (common): adult, fetus, female, male

Taxa Notes

Animals, Chordates, Mammals, Nonhuman Vertebrates, Nonhuman Mammals, Rodents, Vertebrates

RN 127-08-2 (potassium acetate)

L116 ANSWER 2 OF 7 BIOSIS COPYRIGHT (c) 2004 The Thomson Corporation. on STN

AN 2004:181735 BIOSIS

DN PREV200400185021

TI Food additive or product or a pharmaceutical preparation, comprising hydroxymatairesinol.

AU Ahotupa, Markku [Inventor, Reprint Author]; Eckerman, Christer [Inventor]; Kangas, Lauri [Inventor]; Makela, Sari [Inventor]; Saarinen, Niina [Inventor]; Santti, Risto [Inventor]; Warri, Anni [Inventor]

CS Turku, Finland

ASSIGNEE: Hormos Nutraceutical Oy Ltd., Turku, Finland

PI US 6689809 February 10, 2004

Official Gazette of the United States Patent and Trademark Office Patents, (Feb 10 2004) Vol. 1279, No. 2. http://www.uspto.gov/web/menu/patdata.html . e-file.

```
ISSN: 0098-1133 (ISSN print).
DT
     Patent
     English
LA
ED
     Entered STN: 7 Apr 2004
     Last Updated on STN: 7 Apr 2004
AB
     This invention relates to methods for prevention of cancers, certain
     non-cancer, hormone dependent diseases and/or cardiovascular diseases in a
     person, based on administering of hydroxymatairesinol to said
     person. The invention also concerns a method for increasing the level of
     enterolactone or another metabolite of hydroxymatairesinol
     in a person's serum thereby causing prevention of a cancer or a certain
     non-cancer, hormone dependent disease in a person, based on administering
     of hydroxymatairesinol to said person. Furthermore, this
     invention relates to pharmaceutical preparations, food additives and food
     products comprising hydroxymatairesinol.
NCL
     514473000
     Pathology - Therapy 12512
Nutrition - General studies, nutritional status and methods
                                                                    13202
     Food technology - General and methods
                                             13502
     Cardiovascular system - Heart pathology
                                                14506
     Cardiovascular system - Blood vessel pathology
                           17002
     Endocrine - General
     Pharmacology - General
                              22002
     Pharmacology - Cardiovascular system
     Pharmacology - Endocrine system
                                      22016
     Neoplasms - Pathology, clinical aspects and systemic effects
     Neoplasms - Therapeutic agents and therapy
IT
     Major Concepts
        Foods; Pharmacology
IT
     Diseases
        cancer: neoplastic disease
        Neoplasms (MeSH)
IT
     Diseases
        cardiovascular disease: heart disease, vascular disease
        Cardiovascular Diseases (MeSH)
IT
     Diseases
        hormone dependent disease: endocrine disease
IT
     Chemicals & Biochemicals
          enterolactone; hydroxymatairesinol:
        antineoplastic-drug, cardiovascular-drug, hormone-drug, food additive
RN
     78473-71-9 (enterolactone)
       20268-71-7 (hydroxymatairesinol)
L116 ANSWER 3 OF 7 BIOSIS COPYRIGHT (c) 2004 The Thomson Corporation. on STN
     2002:583081 BIOSIS
DN
     PREV200200583081
     USE OF HYDROXYMATAIRESINOL FOR PREVENTION OF CANCERS,
     NON-CANCER, HORMONE DEPENDENT DISEASES AND CARDIOVASCULAR DISEASES BY
     HYDROXYMATAIRESINOL, AND A PHARMACEUTICAL PREPARATION, FOOD
     ADDITIVE AND FOOD PRODUCT COMPRISING HYDROXYMATAIRESINOL.
     Ahotupa, Markku [Inventor, Reprint author]; Eckerman, Chester
     [Inventor]; Kangas, Lauri [Inventor]; Makela, Sari [Inventor];
     Saarinen, Niina [Inventor]; Santti, Risto [Inventor]; Warri, Anni
     [Inventor]
CS
    Turku, Finland
     ASSIGNEE: Hormos Nutraceutical Oy Ltd., Turku, Finland
PI
    US 6451849 September 17, 2002
SO
    Official Gazette of the United States Patent and Trademark Office Patents,
     (Sep. 17, 2002) Vol. 1262, No. 3. http://www.uspto.gov/web/menu/patdata.ht
    ml. e-file.
    CODEN: OGUPE7. ISSN: 0098-1133.
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DT

LA

Patent

English

ED Entered STN: 13 Nov 2002 Last Updated on STN: 13 Nov 2002

This invention relates to methods for prevention of cancers, certain non-cancer, hormone dependent diseases and/or cardiovascular diseases in a person, based on administering of hydroxymatairesinol to said person. The invention also concerns a method for increasing the level of enterolactone or another metabolite of hydroxymatairesinol in a person's serum thereby causing prevention of a cancer or a certain non-cancer, hormone dependent disease in a person, based on administering of hydroxymatairesinol to said person. Furthermore, this invention relates to pharmaceutical preparations, food additives and food products comprising hydroxymatairesinol.

NCL 514473000

CC Pharmacology - General 22002 Neoplasms - Pathology, clinical aspects and systemic effects 24004 Neoplasms - Therapeutic agents and therapy 24008

IT Major Concepts

Oncology (Human Medicine, Medical Sciences); Pharmacology

IT Diseases

cancer: neoplastic disease, drug therapy
Neoplasms (MeSH)

IT Chemicals & Biochemicals

hydroxymatairesinol: antineoplastic-drug

RN 20268-71-7 (hydroxymatairesinol)

L116 ANSWER 4 OF 7 BIOSIS COPYRIGHT (c) 2004 The Thomson Corporation. on STN

AN 2000:422437 BIOSIS

DN PREV200000422437
TI A novel treatment for lupus nephritis: Lignan precursor derived from flax.

AU Clark, W. F. [Reprint author]; Muir, A. D.; Westcott, N. D.; Parbtani, A.

CS Division of Nephrology, London Health Sciences Centre, 375 South Street, London, Ontario, N6A 4G5, Canada

SO Lupus, (2000) Vol. 9, No. 6, pp. 429-436. print. ISSN: 0961-2033.

DT Article

LA English

ED Entered STN: 4 Oct 2000
Last Updated on STN: 8 Jan 2

Last Updated on STN: 8 Jan 2002

AB Background: Flaxseed has renoprotective effects in animal and human lupus nephritis. We have recently extracted the lignan precursor

(secoisolariresinol diglucoside) (SDG) to determine if this more palatable derivative of flaxseed would exert renoprotection similar to the whole flaxseed in the aggressive MRL/lpr lupus mouse model. Methods: 131 MRL/lpr mice were randomly assigned to saline gavage, 600, 1200 and 4800 mug lignan gavage groups. At 7 weeks, 6 animals underwent platelet aggregating factor (PAF) lethal challenge and 40 were studied with urine collection to determine the levels of secoisolariresinol, enterodiol and enterolactone in the gavaged animals. A baseline study of 10 saline gavaged animals took place at 6 weeks. 25 animals in the saline gavage, 600 and 1200 mug lignan groups were studied at 14 and 22 weeks for GFR, spleen lymphocyte S-phase and organ weight studies. Results: Metabolic studies indicated that secoisolariresinol is the major metabolite absorbed and the lowest lignan dose provides a lengthening in survival for the PAF lethal challenge. Body weight, fluid and water intake studies demonstrated that the lignan was well tolerated. Changes in proteinuria, GFR and renal size showed a time- and dose-dependent protection for the lignan precursor. Cervical lymph node size and spleen lymphocyte cells in the S-phase demonstrated modest dose-dependent reductions in the lignan gavaged groups. Conclusion: SDG was converted in the gut to secoisolarizesinol, which was absorbed and well tolerated by the MRL/lpr mice. Renoprotection was evidenced, in a dose-dependent fashion, by a significant delay in the onset of proteinuria with preservation in GFR and renal size. This study suggests that SDG may have

a therapeutic role in lupus nephritis. Bones, joints, fasciae, connective and adipose tissue - Pathology CC Digestive system - Physiology and biochemistry 15506 Urinary system - Pathology Immunology - Immunopathology, tissue immunology 34508 Major Concepts TT Clinical Immunology (Human Medicine, Medical Sciences); Urology (Human Medicine, Medical Sciences) TΤ Parts, Structures, & Systems of Organisms gut: digestive system ΤT Diseases lupus nephritis: connective tissue disease, immune system disease, urologic disease, treatment Lupus Nephritis (MeSH) IT Diseases proteinuria: urologic disease Proteinuria (MeSH) Chemicals & Biochemicals TT lignan precursor: derived from flax IT Miscellaneous Descriptors body weight; glomerular filtration rate ORGN Classifier Hominidae 86215 Super Taxa Primates; Mammalia; Vertebrata; Chordata; Animalia Organism Name human Taxa Notes Animals, Chordates, Humans, Mammals, Primates, Vertebrates ORGN Classifier Muridae 86375 Super Taxa Rodentia; Mammalia; Vertebrata; Chordata; Animalia Organism Name mouse: animal model Taxa Notes Animals, Chordates, Mammals, Nonhuman Vertebrates, Nonhuman Mammals, Rodents, Vertebrates L116 ANSWER 5 OF 7 BIOSIS COPYRIGHT (c) 2004 The Thomson Corporation. on STN 2000:331401 BIOSIS ANDNPREV200000331401 Hydroxymatairesinol, a novel enterolactone precursor TI with antitumor properties from coniferous tree (Picea abies). Saarinen, N. M. [Reprint author]; Warri, A. [Reprint author]; Makela, S. ΑU I. [Reprint author]; Eckerman, C.; Reunanen, M.; Ahotupa, M. [Reprint author]; Salmi, S. M. [Reprint author]; Franke, A. A.; Kangas, L.; Santti, R. [Reprint author] Department of Anatomy and Medical Research Laboratory, Institute of CS Biomedicine, University of Turku, FIN-20520, Turku, Finland Nutrition and Cancer, (2000) Vol. 36, No. 2, pp. 207-216. print. SO CODEN: NUCADQ. ISSN: 0163-5581. DTArticle English LA EDEntered STN: 2 Aug 2000 Last Updated on STN: 7 Jan 2002 The potential for the extraction of the plant lignan AB hydroxymatairesinol (HMR) in large scale from Norway spruce (Picea abies) has given us the opportunity to study the metabolism and biological actions of HMR in animals. HMR, the most abundant single component of spruce lignans, was metabolized to enterolactone (ENL) as the major metabolite in rats after oral administration. The amounts of

urinary ENL increased with the dose of HMR (from 3 to 50 mg/kg), and only

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Article

English

Entered STN: 2 Aug 2000

Last Updated on STN: 7 Jan 2002

minor amounts of unmetabolized HMR isomers and other lignans were found in urine. HMR (15 mg/kg body wt po) given for 51 days decreased the number of growing tumors and increased the proportion of regressing and stabilized tumors in the rat dimethylbenz(a)anthracene-induced mammary tumor model. HMR (50 mg/kg body wt) did not exert estrogenic or antiestrogenic activity in the uterine growth test in immature rats. also showed no antiandrogenic responses in the growth of accessory sex glands in adult male rats. Neither ENL nor enterodiol showed estrogenic or antiestrogenic activity via a classical alpha- or beta-type estrogen receptor-mediated pathway in vitro at <1.0 muM. HMR was an effective antioxidant in vitro. Pharmacognosy and pharmaceutical botany Biochemistry studies - General Biophysics - General 10502 Pharmacology - General 22002 Neoplasms - General 24002 Plant physiology - Chemical constituents 51522 Major Concepts Biochemistry and Molecular Biophysics; Pharmacognosy (Pharmacology); Tumor Biology Diseases cancer: neoplastic disease Neoplasms (MeSH) Chemicals & Biochemicals hydroxymatairesinol: antitumor properties, enterolactone precursor, oral administration, quantitative structure-activity relationships ORGN Classifier Coniferopsida 25102 Super Taxa Gymnospermae; Spermatophyta; Plantae Organism Name Picea abies [Norway spruce]: medicinal plant Taxa Notes Gymnosperms, Plants, Spermatophytes, Vascular Plants ORGN Classifier 86375 Muridae Super Taxa Rodentia; Mammalia; Vertebrata; Chordata; Animalia Organism Name Sprague-Dawley rat: male Taxa Notes Animals, Chordates, Mammals, Nonhuman Vertebrates, Nonhuman Mammals, Rodents, Vertebrates 20268-71-7 (hydroxymatairesinol) L116 ANSWER 6 OF 7 BIOSIS COPYRIGHT (c) 2004 The Thomson Corporation. 2000:331149 BIOSIS PREV200000331149 Isoflavonoids and lignans have different potentials to modulate oxidative genetic damage in human colon cells. Pool-Zobel, Beatrice L. [Reprint author]; Adlercreutz, Herman; Glei, Michael; Liegibel, Ute M.; Sittlingon, Julie; Rowland, Ian; Wahala, Kristiina; Rechkemmer, Gerhard Department of Nutritional Toxicology, Institute for Nutrition, Friedrich Schiller University, Dornburger Strasse 25, 07743, Jena, Germany Carcinogenesis (Oxford), (June, 2000) Vol. 21, No. 6, pp. 1247-1252. print. CODEN: CRNGDP. ISSN: 0143-3334.

Polyphenolic compounds, including isoflavonoids and lignans, have been AB suggested to be chemopreventive on account of antioxidative properties. In this context it is of importance to have knowledge of their ability to reduce oxidative stress within target cells of tumorigenesis. Therefore, we investigated isoflavonoids and lignans for modulation of oxidative genetic damage in mammalian cells. H2O2-induced damage as well as endogenous DNA strand breaks and oxidized bases were determined after 30 min incubation of human colon cells with polyphenols using various modifications of the microgel electrophoresis assay (Comet assay). Enterolactone, a mammalian metabolite of plant lignans, was additionally investigated for modulation of intracellular oxidative stress in NIH 3T3 cells using laser scanning microscopy. In vivo effects of rye crispbread (a source of lignans) were investigated in 12 human volunteers by determining genetic damage in lymphocytes and antioxidant activity in plasma (FRAP assay). Genistein induced DNA breaks in the human tumour cell line HT29 clone 19A (12.5-100 muM). The polyphenols (100 muM) did not reduce damage induced by 150 muM H2O2, indicating that they lacked antioxidative potential. At this concentration enterolactone also had no effect on intracellular oxidative stress induced by 31.25 and 125 muM H2O2. In contrast, enterolactone, dihydrogenistein and formononetin reduced endogenous oxidative DNA damage at 100 muM. Daily ingestion of nine slices (76.5 g/day) of rye crispbread per day (containing 41.8 and 33.0 mug/100 g dry weight secoisolariciresinol and matairesinol, respectively) for 2 weeks did not significantly reduce genetic damage in blood lymphocytes , nor was there a modulation of plasma antioxidant capacity. The moderate effects of high concentrations of the tested compounds on endogenous oxidative DNA damage and failure to prevent H2O2-induced damage are indicative of only marginal protective potential by antioxidant mechanisms. The genotoxic effects of genistein deserve further investigation. 02506 CC Cytology - Animal Cytology - Human 02508 Genetics - Animal 03506 Genetics - Human 03508 Pathology - Therapy 12512 Digestive system - Physiology and biochemistry Pharmacology - Clinical pharmacology 22005 Neoplasms - Pathology, clinical aspects and systemic effects Neoplasms - Therapeutic agents and therapy TT Major Concepts Genetics; Pharmacology; Tumor Biology Parts, Structures, & Systems of Organisms IT colon cells: digestive system, drug-induced oxidative genetic damage modulation, hydrogen peroxide-induced DNA damage, in-vitro culture IT Chemicals & Biochemicals biochanin A: antineoplastic-drug, chemopreventive agent; daidzein: antineoplastic-drug, chemopreventive agent; enterodiol: antineoplastic-drug, chemopreventive agent; enterolactone: antineoplastic-drug, chemopreventive agent; equol: antineoplastic-drug, chemopreventive agent; formononetin: antineoplastic-drug, chemopreventive agent; genestein: antineoplastic-drug, chemopreventive agent ORGN Classifier Hominidae 86215 Super Taxa Primates; Mammalia; Vertebrata; Chordata; Animalia HT-29 cell line: clone 19, drug-induced oxidative genetic damage modulation, human colon cancer cell line, hydrogen peroxide-induced DNA damage, in-vitro model system human: normal subjects

Taxa Notes

```
Animals, Chordates, Humans, Mammals, Primates, Vertebrates
ORGN Classifier
        Muridae
                  86375
     Super Taxa
        Rodentia; Mammalia; Vertebrata; Chordata; Animalia
     Organism Name
        NIH3T3 cell line: drug-induced oxidative genetic damage modulation,
        hydrogen peroxide-induced DNA damage, in-vitro model system, mouse
        fibroblast cell line
     Taxa Notes
        Animals, Chordates, Mammals, Nonhuman Vertebrates, Nonhuman Mammals,
        Rodents, Vertebrates
     491-80-5 (biochanin A)
RN
     486-66-8 (daidzein)
     80226-00-2 (enterodiol)
       78473-71-9 (enterolactone)
     531-95-3 (equol)
     485-72-3 (formononetin)
L116 ANSWER 7 OF 7 BIOSIS COPYRIGHT (c) 2004 The Thomson Corporation. on STN
     1994:452533 BIOSIS
AN
     PREV199497465533
DN
     Natural flavonoids and lignans are potent cytostatic agents against human
TI
     leukemic HL-60 cells.
     Hirano, Toshihiko; Gotoh, Manabu; Oka, Kitaro
ΑU
     Department Clinical Pharmacology, Tokyo College Pharmacy, 1432-1 Hachioji,
CS
     Tokyo 192-03, Japan
     Life Sciences, (1994) Vol. 55, No. 13, pp. 1061-1069.
SO
     CODEN: LIFSAK. ISSN: 0024-3205.
DT
     Article
LA
     English
     Entered STN: 24 Oct 1994
ED
     Last Updated on STN: 16 Dec 1994
     Anti leukemic-cell efficacy of 28 naturally occurring and synthetic
AB
     flavonoids and 11 naturally occurring lignans on human promyelocytic
     leukemic cell line HL-60 were examined using MTT assay methods.
     Differences between anti cell-proliferative activity and cytotoxicity of
     these compounds were compared with those of 4 clinical anti-cancer agents.
     Eight of the 28 flavonoids and 4 of the 11 lignans showed considerable
     suppressive effects on HL-60 cell growth with IC-50s ranging from 10-940
     ng/ml. Among these compounds, genistein, honokiol, machilin A,
     matairesinol, and arctigenin had the strongest effects with IC-50s
     less than 100 ng/ml, which were almost equivalent to the effects of
     current anti-cancer agents. The flavonoid genistein and the lignans,
     however, showed little or no cytotoxicity against HL-60 cells as assessed
     by dye exclusion tests (LC-50s gt 2,900ng/ml), whereas the regular
     anti-cancer agents had potent cytotoxicity. All of the flavonoids and
     lignans, except for machilin A and arctigenin, were less effective against
     growth of human T lymphocytic leukemia cell line MOLT-4. In
     addition, the flavonoid and the lignans showed little or no inhibiting
     activity on mitogen-induced blastogenesis of human peripheral-blood
     lymphocytes. The lignans and genistein were strongly suppressive
     against incorporations of (3H)thymidine, (3H)uridine, and (3H)leucine into
     HL-60 cells. These results showed that some of the naturally occurring
     flavonoids and lignans inhibited HL-60 cell growth with a non-toxic
     mechanism, possibly via cessation of DNA, RNA, and/or protein synthesis of
     the leukemic cells.
                        02508
     Cytology - Human
     Biochemistry studies - General
                                      10060
     Biochemistry studies - Nucleic acids, purines and pyrimidines
     Biochemistry studies - Proteins, peptides and amino acids
     Pathology - Therapy
                          12512
     Metabolism - Proteins, peptides and amino acids
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Metabolism - Nucleic acids, purines and pyrimidines
     Blood - Blood cell studies
                                  15004
     Blood - Blood, lymphatic and reticuloendothelial pathologies
     Blood - Lymphatic tissue and reticuloendothelial system
                                            22005
     Pharmacology - Clinical pharmacology
     Pharmacology - Blood and hematopoietic agents
     Neoplasms - Therapeutic agents and therapy
     Neoplasms - Blood and reticuloendothelial neoplasms
                                                            24010
     Development and Embryology - Morphogenesis
                                                   25508
     Immunology - Immunopathology, tissue immunology
     Plant physiology - Chemical constituents
                                                 51522
     Pharmacognosy and pharmaceutical botany
IT
     Major Concepts
        Blood and Lymphatics (Transport and Circulation); Cell Biology;
        Clinical Endocrinology (Human Medicine, Medical Sciences); Development;
        Hematology (Human Medicine, Medical Sciences); Metabolism; Oncology
        (Human Medicine, Medical Sciences); Pharmacognosy (Pharmacology);
        Pharmacology
IT
     Chemicals & Biochemicals
        GENISTEIN; HONOKIOL; MACHILIN A; MATAIRESINOL; ARCTIGENIN
IT
     Miscellaneous Descriptors
        ANTINEOPLASTIC-DRUG; ARCTIGENIN; CELL GROWTH; CYTOTOXICITY; GENISTEIN;
        HONOKIOL; IMMUNOSUPPRESSION; MACHILIN A; MATAIRESINOL;
        NUCLEIC ACID SYNTHESIS; PERIPHERAL BLOOD LYMPHOCYTE;
        POTENTIAL THERAPEUTIC APPLICATION; PROTEIN SYNTHESIS
ORGN Classifier
        Aristolochiaceae
                           25595
     Super Taxa
        Dicotyledones; Angiospermae; Spermatophyta; Plantae
     Organism Name
        Asarum sieboldi
     Taxa Notes
        Angiosperms, Dicots, Plants, Spermatophytes, Vascular Plants
ORGN Classifier
        Hominidae
                    86215
     Super Taxa
        Primates; Mammalia; Vertebrata; Chordata; Animalia
     Organism Name
        Hominidae
     Taxa Notes
        Animals, Chordates, Humans, Mammals, Primates, Vertebrates
ORGN Classifier
        Lauraceae
                    26245
     Super Taxa
        Dicotyledones; Angiospermae; Spermatophyta; Plantae
     Organism Name
        Machilus thunbergii
     Taxa Notes
        Angiosperms, Dicots, Plants, Spermatophytes, Vascular Plants
ORGN Classifier
        Organisms
                    00500
     Super Taxa
        Organisms
     Organism Name
        Forsythia viridissima
     Taxa Notes
        Organisms
ORGN Classifier
        Pedaliaceae
                      26525
     Super Taxa
        Dicotyledones; Angiospermae; Spermatophyta; Plantae
     Organism Name
        Sesamum indicum
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Taxa Notes Angiosperms, Dicots, Plants, Spermatophytes, Vascular Plants ORGN Classifier Saxifragaceae 26745 Super Taxa Dicotyledones; Angiospermae; Spermatophyta; Plantae Organism Name Saxifragaceae Taxa Notes Angiosperms, Dicots, Plants, Spermatophytes, Vascular Plants RN 446-72-0 (GENISTEIN) 35354-74-6 (HONOKIOL) 110269-50-6 (MACHILIN A) 580-72-3 (MATAIRESINOL) 7770-78-7 (ARCTIGENIN) => => fil wpix FILE 'WPIX' ENTERED AT 14:42:49 ON 30 SEP 2004 COPYRIGHT (C) 2004 THE THOMSON CORPORATION FILE LAST UPDATED: 28 SEP 2004 <20040928/UP> MOST RECENT DERWENT UPDATE: 200462 <200462/DW> DERWENT WORLD PATENTS INDEX SUBSCRIBER FILE, COVERS 1963 TO DATE >>> FOR A COPY OF THE DERWENT WORLD PATENTS INDEX STN USER GUIDE, PLEASE VISIT: http://www.stn-international.de/training center/patents/stn guide.pdf <<< >>> FOR DETAILS OF THE PATENTS COVERED IN CURRENT UPDATES, SEE http://thomsonderwent.com/coverage/latestupdates/ <<< >>> FOR INFORMATION ON ALL DERWENT WORLD PATENTS INDEX USER GUIDES, PLEASE VISIT: http://thomsonderwent.com/support/userguides/ <<< >>> NEW! FAST-ALERTING ACCESS TO NEWLY-PUBLISHED PATENT DOCUMENTATION NOW AVAILABLE IN DERWENT WORLD PATENTS INDEX FIRST VIEW - FILE WPIFV. FOR FURTHER DETAILS: http://www.thomsonderwent.com/dwpifv <<< >>> NEW DISPLAY FORMAT HITSTR ADDED ALLOWING DISPLAY OF HIT STRUCTURES WITHIN THE BIBLIOGRAPHIC DOCUMENT < => d all abeq tech abex L124 ANSWER 1 OF 1 WPIX COPYRIGHT 2004 THE THOMSON CORP on STN AN2003-658464 [62] WPIX DNC C2003-179746 TIInhibiting overactivity of phagocytes or lymphocytes in individual used for treating or preventing an acute ischemia-reperfusion injury or chronic condition, e.g. rheumatoid arthritis, by administering lignan. DC B03 D13 AHOTUPA, M; ERIKSSON, J; KANGAS, L; KOMI, J; KORTE, H; PERALA, M; UNKILA, IN M; PERAELAE, M PA (AHOT-I) AHOTUPA M; (ERIK-I) ERIKSSON J; (KANG-I) KANGAS L; (KOMI-I) KOMI J; (KORT-I) KORTE H; (PERA-I) PERALA M; (UNKI-I) UNKILA M; (HORM-N) HORMOS NUTRACEUTICAL LTD OY 101 CYC PΙ US 2003100514 A1 20030529 (200362)* 10 A61K031-365

A1 20030605 (200362) EN

RW: AT BE BG CH CY CZ DE DK EA EE ES FI FR GB GH GM GR IE IT KE LS LU

A61K031-34

WO 2003045376

ADT

FDT

AB

FS

TECH

ABEX

MC MW MZ NL OA PT SD SE SK SL SZ TR TZ UG ZM ZW W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EC EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ OM PH PL PT RO RU SD SE SG SI SK SL TJ TM TN TR TT TZ UA UG US UZ VC VN YU ZA A1 20030610 (200419) AU 2002365519 A61K031-34 US 2003100514 A1 US 2001-991971 20011126; WO 2003045376 A1 WO 2002-FI936 20021121; AU 2002365519 A1 AU 2002-365519 20021121 AU 2002365519 A1 Based on WO 2003045376 PRAI US 2001-991971 20011126 ICM A61K031-34; A61K031-365 ICS A61K031-05 US2003100514 A UPAB: 20030928 NOVELTY - Overactivity of phagocytes or lymphocytes in an individual is inhibited by administering a lignan. DETAILED DESCRIPTION - Inhibiting overactivity of phagocytes or lymphocytes in an individual by administering a lignan, where the phagocytes are neutrophils and the lignan is hydroxymatairesinol and/or matairesinol; the phagocytes are cells of myeloid origin and the lignan is enterolactone and/or hydroxymatairesinol; or the lymphocytes are T-lymphocytes and the lignan is hydroxymatairesinol, matairesinol, and/or enterolactone. ACTIVITY - Vasotropic; Cardiant; Cerebroprotective; Antibacterial; Immunosuppressive; Antirheumatic; Antiarthritic; Antiinflammatory; Anti-HIV; Antipsoriatic; Antiallergic; Antiparkinsonian; Nootropic; Neuroprotective; Osteopathic; Antidiabetic; Antiarteriosclerotic; Ophthalmological. MECHANISM OF ACTION - None given. USE - The invention is used for inhibiting overactivity of phagocytes or lymphocytes in an individual, thus treating or preventing an acute ischemia- reperfusion injury or a chronic condition. Acute ischemia-reperfusion injury is an injury in myocardial infarction, stroke, transplantation, adult respiratory distress syndrome, ischemic heart disease, or endotoxic or hemorrhagic shock. Chronic condition includes rheumatoid arthritis, allergic conditions including inflammatory bowel disease or an inflammatory condition of the skin, HIV, AIDS, psoriasis, Parkinson's disease, Alzheimer's disease, autoimmune disease, type I or type II diabetes, hypercholesterolemic atherosclerosis, cataract, osteoporosis or amylotrophic lateral sclerosis. (All claimed) ADVANTAGE - The invention decreases the formation of reactive oxygen species. It lowers the risk, prevents or treats other diseases or conditions, which are not due to lipid, DNA, or protein oxidation but which are due to overactive neutrophils. DESCRIPTION OF DRAWING(S) - The figure shows the oxidative burst and myeloperoxidase activity. Dwg. 1/4 CPI AB; GI; DCN CPI: B07-A02A; B10-E04C; B14-A01; B14-A02B1; B14-C03; B14-C06; B14-C09; B14-F01; B14-F02; B14-F07; B14-G02; B14-J01; B14-J01A3; B14-N03; B14-N17C; B14-S04; D03-H01T2 UPTX: 20030928 TECHNOLOGY FOCUS - BIOLOGY - Preferred Component: The phagocytes are cells of myeloid origin, the TNF-alpha release of which is reduced, and the lignan is enterolactone or hydroxymatairesinol. UPTX: 20030928 ADMINISTRATION - Dosage comprises 10-2000 mg/day, preferably 100-600

mg/day for adult persons, and may be administered orally. Oral dosage forms includes powders, granules, capsules, tablets, caplets, lozenges, liquids, elixirs, emulsions, and suspensions.

EXAMPLE - Monocytes were isolated from human peripheral blood mononuclear cells by magnetic sorting. The cells were pre-incubated with the lignans matairesinol and enterolactone (1-100 micro M) or interleukin-10 (IL-10) (100 U/ml) for 24 hours before addition of LPS (1 micro g/ml) into cell culture. After additional 48 hours the levels of TNF from culture supernatants were measured by enzyme linked immunosorbent assays (ELISA). Results showed that matairesinol and enterolactone were effective at concentration of 100 micro M as the positive control, IL-10.

=> d his

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(FILE 'HOME' ENTERED AT 13:25:41 ON 30 SEP 2004)
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L1
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              3 S 580-72-3 OR 20268-71-7 OR 78473-71-9
L2
                E C20H2207/MF
                E C20H22O7/MF
             37 S E3 AND 46.150.18/RID AND OC4/ES AND 3/NR
L3
             26 S L3 AND 3 METHOXY
L4
             26 S L4 AND 4 HYDROXY
L5
             21 S L5 AND FURANONE
L6
                SEL RN 1 6 7 8 10 11 16 20
              8 S E1-E8
L7
              7 S L7 NOT 718614-97-2
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                SEL RN 4 5
L9
              5 S L8 NOT E9-E10
             32 S L3 NOT L9
L10
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1.11
             12 S L11 AND 4 HYDROXY AND 3 METHOXY AND FURANONE
L12
              4 S L12 NOT (D/ELS OR 13C# OR LABELED)
L13
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L14
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L15
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L16
L17
             14 S L2, L9, L13, L16
              9 S L17 AND (?MATAIRESINOL? OR ?ENTEROLACTON?)/CNS
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L19
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L20
                SEL RN
L21
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L22
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L23
L24
            640 S L22, L23
                E AHOTUPA M/AU
L25
             91 S E3-E5
                E ERIKSSON J/AU
            221 S E3-E11, E34-E36
L26
                E KANGAS L/AU
            127 S E3-E5, E8-E11
L27
                E UNKILA M/AU
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48 S E3-E5

12 S E3-E6

E KOMI J/AU

L28

T₁29

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E PERALA M/AU
             21 S E3, E4, E6
L30
                E KORTE H/AU
             23 S E3, E4, E10
L31
                E HORMOS/PA,CS
             27 S E3-E19
L32
             16 S L24 AND L25-L32
L33
                E PHAGOCYTE/CT
           3427 S E3, E12
L34
                E E12+ALL
                E E2+ALL
          32274 S E5+NT
L35
                E NEUTROPHIL/CT
                E E3+ALL
          29239 S E24, E23
L36
                E T CELL/CT
                E E4+ALL
L37
          40180 S E20-E23
          70058 S E19+NT
L38
                E E18+ALL
L39
         169033 S E19, E18+NT
                E MYELOID/CT
                E E11+ALL
           2697 S E2
L40
              4 S L24 AND L34-L40
L41
                E ANIMAL RESPIRATION/CT
           1613 S E3 (L) BURST
L42
                E RESPIRATION, ANIMAL/CT
           1421 S E4
L43
                E REACTIVE OXYGEN/CT
                E E4+ALL
          22365 S E3
L44
              2 S L24 AND L42-L44
L45
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L46
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              1 S L24 AND L46
L47
                E LIGNAN/CT
                E E4+ALL
            356 S L24 AND E2
L48
L49
            356 S L24 AND E2+NT
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L50
              1 S L50 AND L33
L51
L52
              3 S L50 AND L48, L49
              4 S L50-L52
L53
             65 S L20 (L) (THU OR DMA OR PAC OR PKT)/RL
L54
            160 S L24 AND (PHARMACEUT? OR PHARMACOL? OR PATHOL? OR IMMUN?)/SC,S
L55
            164 S L54, L55
L56
              9 S L56 AND L33
L57
              3 S L56 AND L53
L58
L59
              4 S L53, L58
               8 S L57 NOT L59
L60
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L61
              1 S 9003-99-0
     FILE 'HCAPLUS' ENTERED AT 14:15:57 ON 30 SEP 2004
L62
               2 S L61 AND L24
               1 S L24 AND MYELOPEROXIDASE
L63
               6 S L24 AND ?PEROXIDASE?
L64
L65
              6 S L62-L64
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17 S L59, L60, L65
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L69
L70
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                E TRANSPLANTATION/CT
            812 S E3
L71
                E TRANSPLANT/CT
L72
           ·494 S E3
L73
          87407 S E5+OLD, NT, PFT, RT
           5085 S E61
L74
L75
          76222 S E69+OLD, NT, PFT, RT
L76
            812 S E72, E74
                E E3+ALL
                E E2+ALL
          7719 S E7-E16
L77
          35079 S E6+NT
L78
          6674 S E43+NT
L79
          30585 S E42+NT
L80
              5 S L24 AND L71-L80
L81
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L82
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L83
                SEL HIT RN
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L84
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L85
              4 S L84 AND L20
L86
              2 S L84 AND L61, L46
     FILE 'REGISTRY' ENTERED AT 14:24:50 ON 30 SEP 2004
     FILE 'HCAPLUS' ENTERED AT 14:25:17 ON 30 SEP 2004
L87
              3 S L24 AND ?PHAGOCYT?
              2 S L87 NOT L83
L88
     FILE 'MEDLINE' ENTERED AT 14:26:28 ON 30 SEP 2004
L89
            117 S L20
L90
            215 S L23
            231 S L89, L90
L91
              0 S L91 AND ?PHAGOCYT?
L92
                E PHAGOCYTE/CT
                E E29+ALL
L93
              1 S L91 AND E6+NT
              1 S L91 AND NEUTROPHIL
                E MYELOID CELL/CT
                E E8+ALL
L95
              1 S L91 AND E3+NT
               E LYMPHOCYTE/CT
L96
              1 S L91 AND E6+NT
L97
              0 S L91 AND E21+NT
L98
              0 S L91 AND E43+NT
              0 S L91 AND E109+NT
L99
L100
              3 S L91 AND E157+NT
                E T CELL/CT
                E T CELLS/CT
                E E3+ALL
L101
              0 S L91 AND E2+NT
                E E2+ALL
              3 S L91 AND E15+NT
L102
                E REACTIVE OXYGEN/CT
L103
              0 S L91 AND E4+NT
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E E4+ALL
              1 S L91 AND E13+NT
L104
              2 S L91 AND E12+NT
L105
                E RESPIRATION BURST/CT
                E RESPIRATORY BURST/CT
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L106
              6 S L92-L106
L107
              5 S L107 AND PY<=2001
L108
     FILE 'MEDLINE' ENTERED AT 14:32:05 ON 30 SEP 2004
     FILE 'BIOSIS' ENTERED AT 14:32:12 ON 30 SEP 2004
            338 S L24
L109
              4 S L109 AND (AHOTUPA ? OR ERKISSON ? OR KANGAS ? OR UNKILA ? OR
L110
              0 S L109 AND (PHAGOCYT? OR NEUTROPHIL?)
L111
              3 S L109 AND (T CELL OR LYMPHOCYT?)
L112
              0 S L109 AND MYELOID CELL
L113
              0 S L109 AND MYELOID
L114
              0 S L109 AND (HEMATOPO? OR HAEMATOPO?)
L115
L116
              7 S L110, L112
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     FILE 'WPIX' ENTERED AT 14:35:03 ON 30 SEP 2004
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L117
                E ENTERLOCATONE/DCN
                E HYDROXYATAIRESINOL/DCN
                E HYDROXYMATAIRESINOL/DCN
                E MATAIRESINOL/DCN
                E MATAIRESINOL/CN
              2 S E3
L118
                E HYDROXYMATAIRESINOL/CN
L119
              1 S E3
                E ENTERLOCATONE/CN
                E ENTEROLACTONE/DCN
                E ENTEROLACTONE/CN
L120
              2 S E3
L121
              5 S L118-L120
              0 S 3 4 BIS 3 HYDROXY BENZYL DIHYDRO FURAN 2 ONE
L122
              0 S 3 4 BIS 4 HYDROXY 3 METHOXY BENZYL DIHYDRO FURAN 2 ONE
L123
              1 S L117 AND (PHAGOCYT? OR NEUTROPHIL? OR MYELOID? OR T CELL OR L
L124
     FILE 'WPIX' ENTERED AT 14:42:49 ON 30 SEP 2004
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